Refine Search

Search Results -

Term	Documents
FACTOR .	913155
FACTORS	734419
VIII:C	. 455
VIII:CS	0
WILLEBRAND	4385
WILLEBRANDS	276
CONCENTRATE	287953
CONCENTRATES	86190
((FACTOR ADJ VIII:C) SAME (WILLEBRAND ADJ FACTOR) SAME CONCENTRATE).PGPB,USPT,USOC,EPAB,JPAB,DWPI.	14
(FACTOR VIII:C SAME WILLEBRAND FACTOR SAME CONCENTRATE).PGPB,USPT,USOC,EPAB,JPAB,DWPI.	. 14

US Pre-Grant Publication Full-Text Database US Patents Full-Text Database US OCR Full-Text Database

Database:

EPO Abstracts Database
JPO Abstracts Database
Derwent World Patents Index

IBM Technical Disclosure Bulletins

Search:

L16

Recall Text Clear Interrupt

Search History

DATE: Tuesday, August 02, 2005 Printable Copy Create Case

Set
Name
Side by
Side
Side
Set
Count
Name
result set

DB=PGPB, USPT, USOC, EPAB, JPAB, DWPI; THES=ASSIGNEE; PLUR=YES; OP=ADJ

L16 factor VIII:C same Willebrand factor same concentrate

14 <u>L16</u>

Refine Search

<u>L15</u>	(Willebrand factor)	2	<u>L15</u>
<u>L14</u>	process same producing same concentrate same factor VIIIC same Willebrand factor	0	<u>L14</u>
<u>L13</u>	L2 and @py<2002	25	<u>L13</u>
<u>L12</u>	L10 and antibody	1	<u>L12</u>
<u>L11</u>	L10 and RCoF	0	<u>L11</u>
<u>L10</u>	L9 and producing	1	<u>L10</u>
<u>L9</u>	L8 and process	1	<u>L9</u>
<u>L8</u>	L7 and glycine	1	<u>L8</u>
<u>L7</u>	L5 and alkali metal	1	<u>L7</u>
<u>L6</u>	L4 and fractional precipitation	0	<u>L6</u>
<u>L5</u>	L4 and concentrate	3	<u>L5</u>
<u>L4</u>	L3 and L2 and L1	25	<u>L4</u>
<u>L3</u>	L2 and @py<2002	25	<u>L3</u>
<u>L2</u>	L1 and process same producing	60	<u>L2</u>
T.1	factor VIIIC same Willebrand	171	L1

END OF SEARCH HISTORY

side by side	•	<u>Count</u>	Name result set
DB=P	GPB,USPT,USOC,EPAB,JPAB,DWPI; THES=ASSIGNEE; PLUR=YES;	OP=ADJ	
<u>L18</u>	(fractional precipitation) same (factor VIII:C) and (Willebrand factor)	3	<u>L18</u>
<u>L17</u>	(fractional precipitation) same (factor VIII:C) and (Willebrand factor) same metal	1	<u>L17</u>
<u>L16</u>	factor VIII:C same Willebrand factor same concentrate	14	<u>L16</u>
<u>L15</u>	process same producing same concentrate same (factor VIII:C) same (Willebrand factor)	2	<u>L15</u>
<u>L14</u>	process same producing same concentrate same factor VIIIC same Willebrand factor	0	<u>L14</u>
<u>L13</u>	L2 and @py<2002	25	<u>L13</u>
<u>L12</u>	L10 and antibody	1	<u>L12</u>
<u>L11</u>	L10 and RCoF	0	<u>L11</u>
<u>L10</u>	L9 and producing	1	<u>L10</u>
<u>L9</u>	L8 and process	1	<u>L9</u>
<u>L8</u>	L7 and glycine	1	<u>L8</u>
<u>L7</u>	L5 and alkali metal	1	<u>L7</u>
<u>L6</u>	L4 and fractional precipitation	0	<u>L6</u>
<u>L5</u>	L4 and concentrate	3	<u>L5</u>
<u>L4</u>	L3 and L2 and L1	25	<u>L4</u>
<u>L3</u>	L2 and @py<2002	25	<u>L3</u>
<u>L2</u>	L1 and process same producing	60	<u>L2</u>
I.1	factor VIIIC same Willebrand	171	L1

END OF SEARCH HISTORY

STN

FILE 'RDISCLOSURE' ENTERED AT 11:05:21 ON 02 AUG 2005 COPYRIGHT (C) 2005 Kenneth Mason Publications Ltd.

FILE 'SCISEARCH' ENTERED AT 11:05:21 ON 02 AUG 2005 Copyright (c) 2005 The Thomson Corporation

FILE 'SYNTHLINE' ENTERED AT 11:05:21 ON 02 AUG 2005 COPYRIGHT (C) 2005 Prous Science

FILE 'TOXCENTER' ENTERED AT 11:05:21 ON 02 AUG 2005 COPYRIGHT (C) 2005 ACS

FILE 'USPATFULL' ENTERED AT 11:05:21 ON 02 AUG 2005
CA INDEXING COPYRIGHT (C) 2005 AMERICAN CHEMICAL SOCIETY (ACS)

FILE 'USPAT2' ENTERED AT 11:05:21 ON 02 AUG 2005 CA INDEXING COPYRIGHT (C) 2005 AMERICAN CHEMICAL SOCIETY (ACS)

FILE 'VETB' ENTERED AT 11:05:21 ON 02 AUG 2005 COPYRIGHT (C) 2005 THE THOMSON CORPORATION

FILE 'VETU' ENTERED AT 11:05:21 ON 02 AUG 2005 COPYRIGHT (C) 2005 THE THOMSON CORPORATION

FILE 'WATER' ENTERED AT 11:05:21 ON 02 AUG 2005 COPYRIGHT (C) 2005 Cambridge Scientific Abstracts (CSA)

FILE 'WPIDS' ENTERED AT 11:05:21 ON 02 AUG 2005 COPYRIGHT (C) 2005 THE THOMSON CORPORATION

FILE 'WPIFV' ENTERED AT 11:05:21 ON 02 AUG 2005 COPYRIGHT (C) 2005 THE THOMSON CORPORATION

FILE 'WPINDEX' ACCESS NOT AUTHORIZED

- => s factor VIII:C and Willebrand factor and concentrate and preparation
 - 12 FILES SEARCHED...
 - 21 FILES SEARCHED....
 - 29 FILES SEARCHED...
 - 32 FILES SEARCHED...
 - 47 FILES SEARCHED...
 - 59 FILES SEARCHED...
 70 FILES SEARCHED...
- L1 98 FACTOR VIII:C AND WILLEBRAND FACTOR AND CONCENTRATE AND PREPARAT
- => s L1 and py<2002
 - 1 FILES SEARCHED...
- '2002' NOT A VALID FIELD CODE
 - 4 FILES SEARCHED...
 - 9 FILES SEARCHED...
 - 12 FILES SEARCHED...
 - 15 FILES SEARCHED...
 - 16 FILES SEARCHED...
- 20 FILES SEARCHED...
- '2002' NOT A VALID FIELD CODE
- 29 FILES SEARCHED...
- '2002' NOT A VALID FIELD CODE
- 34 FILES SEARCHED...
- '2002' NOT A VALID FIELD CODE
 - 38 FILES SEARCHED...
 - 44 FILES SEARCHED...
 - 47 FILES SEARCHED...
 - 52 FILES SEARCHED...
- '2002' NOT A VALID FIELD CODE
- '2002' NOT A VALID FIELD CODE
 - 60 FILES SEARCHED...

62 FILES SEARCHED... 68 FILES SEARCHED... 70 FILES SEARCHED...

L2 77 L1 AND PY<2002

=> dup rem

ENTER L# LIST OR (END):L2

DUPLICATE IS NOT AVAILABLE IN 'ADISINSIGHT, ADISNEWS, BIOCOMMERCE, DGENE, DRUGMONOG2, FEDRIP, FOREGE, GENBANK, IMSPRODUCT, IMSRESEARCH, KOSMET, NUTRACEUT, PCTGEN, PHAR, PHARMAML, PROUSDDR, PS, RDISCLOSURE, SYNTHLINE'.

ANSWERS FROM THESE FILES WILL BE CONSIDERED UNIQUE

PROCESSING COMPLETED FOR L2

72 DUP REM L2 (5 DUPLICATES REMOVED)

=> d his

L1

(FILE 'HOME' ENTERED AT 11:04:28 ON 02 AUG 2005)

FILE 'ADISCTI, ADISINSIGHT, ADISNEWS, AGRICOLA, ANABSTR, ANTE, AQUALINE, AQUASCI, BIOBUSINESS, BIOCOMMERCE, BIOENG, BIOSIS, BIOTECHDS, BIOTECHNO, CABA, CANCERLIT, CAPLUS, CEABA-VTB, CEN, CIN, CONFSCI, CROPB, CROPU, DDFB, DGENE, DISSABS, DRUGB, DRUGMONOG2, ...' ENTERED AT 11:05:21 ON 02 AUG 2005

98 S FACTOR VIII:C AND WILLEBRAND FACTOR AND CONCENTRATE AND PREPA

L2 77 S L1 AND PY<2002

L3 72 DUP REM L2 (5 DUPLICATES REMOVED)

=> d L3 1-72 ibib, abs

L3 ANSWER 1 OF 72 USPATFULL on STN

ACCESSION NUMBER:

2004:317310 USPATFULL

TITLE:

Method for purifying factor VIII/vWF company by

cation-exchange chromatography

INVENTOR(S):

Mitterer, Artur, Mannsdorf, AUSTRIA Fischer, Bernhard, Vienna, AUSTRIA Schonberger, Oyvind L., Vienna, AUSTRIA

Thomas-Urban, Kathrin, Freiburg, GERMANY, FEDERAL

REPUBLIC OF

Dorner, Friedrich, Vienna, GERMANY, FEDERAL REPUBLIC OF

Eibl, Johann, Vienna, AUSTRIA

PATENT ASSIGNEE(S):

Baxter Aktiengesellschaft, Vienna, AUSTRALIA (non-U.S.

corporation)

	NUMBER	KIND	DATE		
PATENT INFORMATION:	US 6831159	B1	20041214		
	WO 9838220		19980903		<
APPLICATION INFO.:	US 2000-367459		20000508	(9)	
	WO 1998-AT43		19980227		
			20000508	PCT 371 c	date

NUMBER · DATE

PRIORITY INFORMATION: AT 1997-338 19970227

DOCUMENT TYPE: Utility

FILE SEGMENT: GRANTED PRIMARY EXAMINER: Low, Chris

Low, Christopher S. F.

ASSISTANT EXAMINER: Robinson, Hope A.

LEGAL REPRESENTATIVE: Townsend and Townsend and Crew LLP

NUMBER OF CLAIMS: 11 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS:

0 Drawing Figure(s); 2 Drawing Page(s)

LINE COUNT: 656

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB Methods for recovering factor VIII/vWF-complex that involve binding factor VIII/vWF-complex from a protein solution to a cation exchanger and recovering factor VIII/vWF-complex by step-wise elution are disclosed. The recovered complex contains high-molecular vWF multimers.

Compositions containing factor VIII/vWF-complex as purified by cation exchange chromatography are also provided.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 2 OF 72 USPATFULL on STN

ACCESSION NUMBER: 2003:161896 USPATFULL

Method of recovering highly purified vWF or factor TITLE:

VIII/vWF-complex

Mitterer, Artur, Mannsdorf, AUSTRIA INVENTOR(S):

> Fiedler, Christian, Vienna, AUSTRIA Fischer, Bernhard, Vienna, AUSTRIA Dorner, Friedrich, Vienna, AUSTRIA

Eibl, Johann, Vienna, AUSTRIA

Baxter Aktiengesellschaft, Vienna, AUSTRIA (non-U.S. PATENT ASSIGNEE(S):

corporation)

NUMBER KIND DATE ______ US 6579723 B1 20030617 WO 9838218 19980903 PATENT INFORMATION:

<--

US 1999-367362 APPLICATION INFO.: 19991021 (9) WO 1998-AT33 19980218

> DATE NUMBER

PRIORITY INFORMATION: AT 1997-339 19970227

Utility DOCUMENT TYPE: GRANTED FILE SEGMENT: Le, Long V. PRIMARY EXAMINER:

ASSISTANT EXAMINER: Gabel, Gailene R.

LEGAL REPRESENTATIVE: Townsend and Townsend and Crew LLP

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

NUMBER OF DRAWINGS: 1 Drawing Figure(s); 1 Drawing Page(s)

LINE COUNT: 1046

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

A method for purifying factor VIII/vWF complex or free vWF by AB immunoaffinity chromatography in a form suitable for use as a medicament. Factor VIII/vWF complex or free vWF is recovered from an immunoaffinity adsorbent by using an eluting agent containing a zwitterionic species. The presence of the zwitterionic species allows for the use of mild conditions throughout the preparation, facilitating retention of molecular integrity, activity, and incorporation of the recovered proteins into pharmaceutical preparations without the need for additional stabilizers or preservatives.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 3 OF 72 USPATFULL on STN 1.3

INVENTOR(S):

ACCESSION NUMBER: 2002:268871 USPATFULL

TITLE: Purification of von Willebrand factor by cation exchanger chromatography

Fischer, Bernhard, Vienna, AUSTRIA Schonberger, Oyvind L., Vienna, AUSTRIA Mitterer, Artur, Mannsdorf, AUSTRIA

Fiedler, Christian, Vienna, AUSTRIA Dorner, Friedrich, Vienna, AUSTRIA

Eibl, Johann, Vienna, AUSTRIA

Baxter Aktiengesellschaft, Vienna, AUSTRALIA (non-U.S. PATENT ASSIGNEE(S):

corporation)

NUMBER	KIND	DATE	
 US 6465624 WO 9838219	B1	20021015	
us 1999-367460		19991021	(9)

19980218

19991021 PCT 371 date

NUMBER DATE ______

PRIORITY INFORMATION: AT 1997-337 19970227

DOCUMENT TYPE: Utility FILE SEGMENT: GRANTED

PRIMARY EXAMINER: Carlson, Karen Cochrane Robinson, Hope A.

ASSISTANT EXAMINER:

LEGAL REPRESENTATIVE: Townsend and Townsend and Crew LLP

NUMBER OF CLAIMS: 22 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 3 Drawing Figure(s); 3 Drawing Page(s)

726 LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Disclosed are a method of recovering vWF in which vWF at a low salt concentration is bound to a cation exchanger and vWF having a high specific activity is recovered by fractionated elution, as well as a preparation having purified vWF obtainable by this method.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 4 OF 72 USPATFULL on STN

2002:160854 USPATFULL ACCESSION NUMBER:

Method of chromatographically purifying or TITLE: fractionating, respectively, von Willebrand

factor from a VWF-containing starting material

INVENTOR(S): Siekmann, Juergen, Vienna, AUSTRIA

Turecek, Peter, Klosterneuburg, AUSTRIA Schwarz, Hans-Peter, Vienna, AUSTRIA

Eibl, Johann, Vienna, AUSTRIA

Fischer, Bernhard, Vienna, AUSTRIA Mitterer, Artur, Mannsdorf, AUSTRIA Dorner, Friedrich, Vienna, AUSTRIA

Baxter Aktiengesellschaft, Vienna, AUSTRIA (non-U.S. PATENT ASSIGNEE(S):

corporation)

NUMBER KIND DATE ______ US 6414125 B1 20020702 WO 9833820 19980806 US 1999-355865 19991021 WO 1998-AT20 19980130 PATENT INFORMATION: 19960001 19991021 (9) APPLICATION INFO.: WO 1998-AT20 19980130

19991021 PCT 371 date

NUMBER DATE _____

PRIORITY INFORMATION: AT 1997-176 19970204

DOCUMENT TYPE: Utility GRANTED FILE SEGMENT:

ASSISTANT EXAMINER: Carlson, Karen Cochrane
Robinson Hora LEGAL REPRESENTATIVE: Townsend and Townsend and Crew LLP

NUMBER OF CLAIMS: 47 EXEMPLARY CLAIM:

NUMBER OF DRAWINGS: 1 Drawing Figure(s); 1 Drawing Page(s)

LINE COUNT: 659

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Disclosed is a method of chromatographically purifying or fractionating, respectively, von Willebrand factor (vWF) from a vWF-containing starting material, comprising the following steps:

adsorbing the vWF from the starting material on avid collagen immobilized on a carrier,

separating the non-adsorbed portion and, optionally, washing the carrier,

eluting the vWF from immobilized collagen, and

recovering the purified vWF, as well as a pharmaceutical preparation comprising biologically active vWF which is bound to collagen in a stable manner.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 5 OF 72 USPATFULL on STN

ACCESSION NUMBER: 2002:57411 USPATFULL

Immunotolerant prothrombin complex preparation TITLE:

INVENTOR(S): Schwarz, Hans-Peter, Vienna, AUSTRIA

Turecek, Peter, Klosterneuburg, AUSTRIA

Baxter Aktiengesellschaft, Vienna, AUSTRIA (non-U.S. PATENT ASSIGNEE(S):

corporation)

	NUMBER	KIND	DATE			
PATENT INFORMATION:	US 6358534	B1	20020319			
	WO 9844942		19981015			
APPLICATION INFO .:	US 2000-402582		20000128	(9)		
	WO 1998-AT91		19980406			
			20000128	PCT	371	date

		NUMBER	DATE
PRIORITY	INFORMATION:	 1997-594 1997-1592	19970408 19970919

DOCUMENT TYPE: Utility GRANTED FILE SEGMENT: Witz, Jean C. PRIMARY EXAMINER:

Oppenheimer Wolff & Donnelly LLP LEGAL REPRESENTATIVE:

NUMBER OF CLAIMS: 60 EXEMPLARY CLAIM: 1

0 Drawing Figure(s); 0 Drawing Page(s) NUMBER OF DRAWINGS:

LINE COUNT: 928

The invention relates to an immunotolerant prothrombin complex AB

preparation, a method of producing this preparation, as well as the use of the preparation for producing a

medicament,

ANSWER 6 OF 72 BIOSIS COPYRIGHT (c) 2005 The Thomson Corporation on STN L3

DUPLICATE 1

ACCESSION NUMBER: 2001:524942 BIOSIS PREV200100524942 DOCUMENT NUMBER:

Pasteurized, purified von Willebrand TITLE: factor concentrate and a process for the

preparation thereof.

Heimburger, Norbert [Inventor, Reprint author]; Kumpe, AUTHOR(S):

Gerhard [Inventor]; Wellner, Klaus [Inventor]

Marburg, Germany CORPORATE SOURCE:

ASSIGNEE: Aventis Behring GmbH, Marburg, Germany

PATENT INFORMATION: US 6239261 20010529

Official Gazette of the United States Patent and Trademark SOURCE:

Office Patents, (May 29, 2001) Vol. 1246, No. 5. e-file.

CODEN: OGUPE7. ISSN: 0098-1133.

DOCUMENT TYPE: Patent English LANGUAGE:

ENTRY DATE: Entered STN: 14 Nov 2001

Last Updated on STN: 23 Feb 2002

A process for the preparation of a concentrate of von Willebrand factor is described, entailing a solution of

a complex of this factor with factor VIII:C

being optionally pasteurized and treated with an anion exchanger, there

being no binding of the von Willebrand factor.

ANSWER 7 OF 72 USPATFULL on STN L3

ACCESSION NUMBER: 2001:67424 USPATFULL

TITLE: Stable factor VIII/von Willebrand

factor complex

INVENTOR(S): Fischer, Bernhard, Vienna, Austria

> Mitterer, Artur, Mannsdorf, Austria Dorner, Friedrich, Vienna, Austria

> > KIND DATE

Eibl, Johann, Vienna, Austria

Baxter Aktiengesellschaft, Vienna, Austria (non-U.S. PATENT ASSIGNEE(S):

corporation)

US 6228613 WO 9734930 B1 20010508 <--PATENT INFORMATION: 19970925

NUMBER

US 1998-142768 WO 1997-AT55 19981106 (9) APPLICATION INFO.:

19970313

19981106 PCT 371 date 19981106 PCT 102(e) date

DATE NUMBER -----

PRIORITY INFORMATION: AT 1996-494 19960315

DOCUMENT TYPE: Utility

FILE SEGMENT: Granted
PRIMARY EXAMINER: Carlson, Karen Cochrane
ASSISTANT EXAMINER: Robinson, Hope A.
LEGAL REPRESENTATIVE: Heller Ehrman White & McAuliffe

NUMBER OF CLAIMS: 40 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 9 Drawing Figure(s); 9 Drawing Page(s)

1098 LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

There are disclosed a stable factor VIII/vWF-complex, particularly comprising high-molecular vWF multimers, being free from low-molecular vWF molecules and from proteolytic vWF degradation products, as well as

a method of producing this complex.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L3 ANSWER 8 OF 72 USPATFULL on STN

ACCESSION NUMBER: 2001:63240 USPATFULL

TITLE: Pharmaceutical preparation for treating blood

coagulation disorders

INVENTOR(S): Turecek, Peter, Klosterneuburg/Weidling, Austria

Schwarz, Hans-Peter, Vienna, Austria

Eibl, Johann, Vienna, Austria

Baxter Aktiengesellschaft, Vienna, Austria (non-U.S. PATENT ASSIGNEE(S):

corporation)

NUMBER KIND DATE ______

PATENT INFORMATION: US 6224862 B1 20010501 APPLICATION INFO.: US 2000-521219 20000308 (9)

Division of Ser. No. US 1999-245339, filed on 5 Feb RELATED APPLN. INFO.:

1999 Division of Ser. No. US 1998-165745, filed on 6 Oct 1998, now patented, Pat. No. US 6039945 Division of Ser. No. US 1997-821763, filed on 20 Mar 1997, now

patented, Pat. No. US 5866122, issued on 2 Feb 1999

NUMBER DATE ______ AT 1996-518 19960320 AT 1996-1573 19960904 PRIORITY INFORMATION: AT 1996-1573 19960920

AT 1996-1673 DOCUMENT TYPE: FILE SEGMENT: Utility Granted

PRIMARY EXAMINER: Weddington, Kevin E. LEGAL REPRESENTATIVE: Heller Ehrman White & McAuliffe

NUMBER OF CLAIMS: 4 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 10 Drawing Figure(s); 8 Drawing Page(s)

LINE COUNT: 1454

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

There is disclosed a pharmaceutical preparation for treating

blood coagulation disorders which comprises purified prothrombinase factors, in particular purified prothrombin and optionally purified

factor Xa as active component.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 9 OF 72 USPATFULL on STN

200<u>1:482</u>16 USPATFULL ACCESSION NUMBER:

Process for preparing factor V-deficient plasma, and a TITLE:

deficient plasma which is obtained in this way

Kraus, Michael, Marburg, Germany, Federal Republic of INVENTOR(S):

Aillaud, Erika, Rauschenberg, Germany, Federal Republic

Drescher, Heinz-Hermann, Neustadt, Germany, Federal

Republic of

Dade Behring Marburg GmbH, Marburg, Germany, Federal PATENT ASSIGNEE(S):

Republic of (non-U.S. corporation)

KIND DATE NUMBER ______

US 6211344 B1 20010403 PATENT INFORMATION:

US 1997-915585 APPLICATION INFO.: 19970821 (8)

> NUMBER DATE _____

PRIORITY INFORMATION: DE 1996-19634312 19960824

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: Saunders, David ASSISTANT EXAMINER: De Cloux, Amy

LEGAL REPRESENTATIVE: Finnegan, Henderson, Farabow, Garrett & Dunner, L.L.P.

18 NUMBER OF CLAIMS: EXEMPLARY CLAIM: 1 LINE COUNT: 457

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The invention relates to a process for preparing factor V-deficient plasma, in particular a factor V-deficient plasma from a starting plasma using antibodies, and a deficient plasma which is obtained in this way.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 10 OF 72 USPATFULL on STN L3

2001:26019 USPATFULL ACCESSION NUMBER:

Recombinant factor VIII binding peptides TITLE: Chen, Li Ang, Waverly, TN, United States INVENTOR(S):

Buettner, Joseph A., Raleigh, NC, United States Carbonell, Ruben G., Raleigh, NC, United States

Bayer Corporation, Berkeley, CA, United States (U.S. PATENT ASSIGNEE(S):

corporation)

NUMBER KIND DATE

US 6191256 B1 20010220 US 1998-196934 19981120 PATENT INFORMATION:

19981120 (9) APPLICATION INFO.:

DOCUMENT TYPE: Utility Granted FILE SEGMENT:

Low, Christopher S. F. PRIMARY EXAMINER: ASSISTANT EXAMINER: Mohamed, Abdel A.

Beck, Michael J., Giblin, James A. LEGAL REPRESENTATIVE:

NUMBER OF CLAIMS: EXEMPLARY CLAIM: 1 LINE COUNT: 627

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Peptides that have domains that bind to recombinant factor VIII (rFVIII) are disclosed. A method of rFVIII binding assay using the peptides deduced from a combinatorial library in a filtration plate process is described. A method of using peptides having these available binding domains in an affinity chromatography process to purify factor VIII is also taught.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 11 OF 72 USPATFULL on STN

ACCESSION NUMBER: 2000:174602 USPATFULL

TITLE: Pharmaceutical preparation for treating blood

coagulation disorders

Turecek, Peter, Klosterneuburg/Weidling, Austria INVENTOR(S):

Schwarz, Hans-Peter, Vienna, Austria

Eibl, Johann, Vienna, Austria

Baxter Aktiengesellschaft, Vienna, Austria (non-U.S. PATENT ASSIGNEE(S):

corporation)

KIND DATE NUMBER

20001226 US 6165974 PATENT INFORMATION:

US 1999-245339 19990205 (9) APPLICATION INFO.:

Division of Ser. No. US 1998-165745, filed on 6 Oct RELATED APPLN. INFO.: 1998, now patented, Pat. No. US 6039945 which is a

division of Ser. No. US 1997-821763, filed on 20 Mar 1997, now patented, Pat. No. US 5866122, issued on 2

Feb 1999

NUMBER

PRIORITY INFORMATION: AT 1996-518 19960320 AT 1996-1573 19960904 AT 1996-1673 19960920

Utility DOCUMENT TYPE: FILE SEGMENT: Granted

PRIMARY EXAMINER: Weddington, Kevin E.

LEGAL REPRESENTATIVE: Foley & Lardner

NUMBER OF CLAIMS: 19 EXEMPLARY CLAIM:

NUMBER OF DRAWINGS: 10 Drawing Figure(s); 8 Drawing Page(s)

LINE COUNT: 1552

INVENTOR(S):

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

There is disclosed a pharmaceutical preparation for treating

blood coagulation disorders which comprises purified prothrombinase factors, in particular purified prothrombin and optionally purified

factor Xa as active component.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 12 OF 72 USPATFULL on STN

~2000:105877 USPATFULL ACCESSION NUMBER:

Method for isolation of highly pure von TITLE:

willebrand factor

Fischer, Bernhard, Vienna, Austria Mitterer, Artur, Orth/Donau, Austria Dorner, Friedrich, Vienna, Austria Schwarz, Hans-Peter, Vienna, Austria

Turecek, Peter, Vienna, Austria Eibl, Johann, Vienna, Austria

Falkner, Falko-Guenter, Orth/Donau, Austria

Schlokat, Uwe, Orth/Donau, Austria Mundt, Wolfgang, Vienna, Austria Reiter, Manfred, Vienna, Austria

Den-Bouwmeester, Renate, Vienna, Austria

Immuno Aktiengesellschaft, Vienna, Austria (non-U.S. PATENT ASSIGNEE(S):

	corporation)			
	NUMBER			
PATENT INFORMATION: APPLICATION INFQ.: RELATED APPLN. INFO.:	US 1997-898130 Division of Ser. 1996, now patente	No. US d, Pat.	20000815 19970722 1996-6532 No. US	< (8) 298, filed on 24 May 5854403 which is a -EP3892, filed on 2 Oct
	NUMBER		E	
PRIORITY INFORMATION:	DE 1994-4435485 WO 1995-EP3892	19941		
DOCUMENT TYPE: FILE SEGMENT: PRIMARY EXAMINER: LEGAL REPRESENTATIVE: NUMBER OF CLAIMS: EXEMPLARY CLAIM: NUMBER OF DRAWINGS:	Granted Patterson, Jr., C Foley & Lardner 13 1 1 Drawing Figure(age(s)
LINE COUNT:	793			
CAS INDEXING IS AVAILABLE FOR THIS PATENT. AB The invention relates to a method for isolation of highly pure von Willebrand Factor in which recombinant von Willebrand Factor (rvWF) is chromatographically purified by anion exchange chromatography on an anion exchanger of the quaternary amino type in a buffer solution comprising buffer substances and optionally salt.				
other additives.	According to this	method	, highly	lizers, amino acids and pure recombinant vWF proteins, especially

free from Factor VIII, and is physiologically active.

Further, the invention relates to a pharmaceutical preparation that contains rvWF, which is comprised of multimers with a high structural integrity.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L3	ANSWER	13	OF	72	USPATFULL	on	\mathtt{STN}

ACCESSION NUMBER:

2000:101870 USPATFULL Pharmaceutical preparation for treating blood TITLE:

coagulation disorders

Turecek, Peter, Klosterneuburg/Weidling, Austria Schwarz, Hans-Peter, Vienna, Austria INVENTOR(S):

KIND

DATE

Eibl, Johann, Vienna, Austria

PATENT ASSIGNEE(S): Baxter Aktiengesellschaft, Vienna, Austria (non-U.S.

corporation)

NUMBER

	WOLLDER	11111	21.11	
PATENT INFORMATION:	US 6099837		20000808	<
APPLICATION INFO.:	US 1999-244762		19990205	(9)
RELATED APPLN. INFO.:	Division of Ser.	No. US	1998-1657	45, filed on 6 Oct
	1998 which is a	division	n of Ser.	No. US 1997-821763,
	filed on 20 Mar	1997, no	ow patente	d, Pat. No. US 5866122

	NUMBER	DATE
PRIORITY INFORMATION:	AT 1996-518 AT 1996-1573	19960320 19960904
	AT 1996-1673	19960920
DOCUMENT TYPE:	Utility Granted	

PRIMARY EXAMINER: Weddington, Kevin E. LEGAL REPRESENTATIVE: Foley & Lardner

NUMBER OF CLAIMS: EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 8 Drawing Figure(s); 8 Drawing Page(s)

LINE COUNT: 1533

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

There is disclosed a pharmaceutical preparation for treating

blood coagulation disorders which comprises purified prothrombinase factors, in particular purified prothrombin and optionally purified

factor Xa as active component.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 14 OF 72 USPATFULL on STN

ACCESSION NUMBER: 2000:34192 USPATFULL

Pharmaceutical preparation for treating blood TITLE:

coaqulation disorders

Turecek, Peter, Klosterneuburg/Weidling, Austria INVENTOR(S):

Schwarz, Hans-Peter, Vienna, Austria

Eibl, Johann, Vienna, Austria

PATENT ASSIGNEE(S): Baxter Aktiengesellschaft, Vienna, Austria (non-U.S.

corporation)

NUMBER KIND DATE _____

US 6039945 20000321 US 1998-165745 19981006 (9) PATENT INFORMATION:

APPLICATION INFO.:

RELATED APPLN. INFO.: Division of Ser. No. US 1997-821763, filed on 20 Mar

1997, now patented, Pat. No. US 5866122

NUMBER DATE -----

AT 1996-518 19960320 AT 1996-1573 19960904 AT 1996-1673 19960920 PRIORITY INFORMATION:

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: Weddington, Kevin E. LEGAL REPRESENTATIVE: Foley & Lardner

13 NUMBER OF CLAIMS: EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 8 Drawing Figure(s); 8 Drawing Page(s)

LINE COUNT: 1524

INVENTOR(S):

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

There is disclosed a pharmaceutical preparation for treating

blood coagulation disorders which comprises purified prothrombinase factors, in particular purified prothrombin and optionally purified

factor Xa as active component.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 15 OF 72 USPATFULL on STN

1999:75764 USPATFULL ACCESSION NUMBER:

Protein formulation comprising coagulation factor VIII TITLE:

> or factor IX in an aqueous solution Osterberg, Thomas, Stockholm, Sweden Fatouros, Angelica, Stockholm, Sweden

PATENT ASSIGNEE(S): Pharmacia & Upjohn AB, Stockholm, Sweden (non-U.S.

corporation)

NUMBER KIND DATE US 5919908 19990706 <--PATENT INFORMATION: WO 9630041 . 19961003 <--APPLICATION INFO .: US 1997-913263 19971126 (8) WO 1996-SE419 19960309 19971126 PCT 371 date 19971126 PCT 102(e) date

NUMBER DATE _____

SE 1995-1189 19950331 PRIORITY INFORMATION:

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: Tsang, Cecilia J. Mohamed, Abdel A. ASSISTANT EXAMINER: LEGAL REPRESENTATIVE: Dinsmore & Shohl LLP

NUMBER OF CLAIMS: EXEMPLARY CLAIM: 1 791 LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

A final drug product comprises a plasma protein selected from the group consisting of coagulation factor VIII and factor IX, in an aqueous solution. The concentration of oxygen in the solution is reduced and/or the solution contains an antioxidant. The solution further contains a carbohydrate in a concentration of at least 350 mg/ml. The protein activity after storage for at least 6 months at a temperature of from 0° C. to 40° C. is from 70% to 130% of its initial value. In a process for preparing the final drug product and a method for improving the long-term stability of coagulation factor VIII or factor IX in an aqueous solution, a carbohydrate is included in the solution in a concentration of at least 350 mg/ml and the solution is stored in its final container under an oxygen-reduced atmosphere or includes an antioxidant.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 16 OF 72 USPATFULL on STN

1999:75622 USPATFULL ACCESSION NUMBER:

Composition comprising coagulation factor VIII TITLE:

formulation, process for its preparation and

use of a surfactant as stabilizer Osterberg, Thomas, Stockholm, Sweden

INVENTOR(S): Fatouros, Angelica, Stockholm, Sweden

Pharmacia & Upjohn Aktiebolag, Stockholm, Sweden PATENT ASSIGNEE(S):

(non-U.S. corporation)

NUMBER KIND DATE _____

PATENT INFORMATION: US 5919766 19990706 APPLICATION INFO.: US 1997-863198 19970527 (8)

RELATED APPLN. INFO.: Continuation of Ser. No. US 1994-244266, filed on 20

May 1994, now patented, Pat. No. US 5733873

NUMBER DATE _____ SE 1992-2878 19921002 SE 1993-1580 19930507 SE 1993-2006 19930611 PRIORITY INFORMATION: DOCUMENT TYPE: Utility

FILE SEGMENT: Granted
PRIMARY EXAMINER: Degen, Nancy

LEGAL REPRESENTATIVE: Pollock, Vande Sande & Amernick

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

NUMBER OF DRAWINGS: 2 Drawing Figure(s); 1 Drawing Page(s)

LINE COUNT: 618

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The present invention relates to a novel composition comprising AB coagulation Factor VIII and a non-ionic surfactant such as block co-polymers, e.g. polyoxamers or polyoxyethylene (20) fatty acid esters e.g. polysorbate 20 or polysorbate 80 as a stabilizer. The composition can also comprise sodium chloride, calcium chloride, L-histidine and/or sugars or sugar alcohols.

ANSWER 17 OF 72 USPATFULL on STN T.3

ACCESSION NUMBER: 1999:30944 USPATFULL

TITLE: Method for isolation of highly pure von

willebrand factor

INVENTOR(S): Fischer, Bernhard, Vienna, Austria

> Mitterer, Artur, Orth/Donau, Austria Dorner, Friedrich, Vienna, Austria .Schwarz, Hans-Peter, Vienna, Austria

Turecek, Peter, Vienna, Austria Eibl, Johann, Vienna, Austria

Falkner, Falko-Guenter, Orth/Donau, Austria

<--

Schlokat, Uwe, Orth/Donau, Austria Mundt, Wolfgang, Vienna, Austria Reiter, Manfred, Vienna, Austria

Den-Bouwmeester, Renate, Vienna, Austria

Immuno Aktiengesellschaft, Vienna, Austria (non-U.S. PATENT ASSIGNEE(S):

corporation)

KIND DATE NUMBER

US 1997-898129 19970722 PATENT INFORMATION:

APPLICATION INFO.: 19970722 (8)

Division of Ser. No. US 1996-653298, filed on 24 May RELATED APPLN. INFO.:

NUMBER DATE

PRIORITY INFORMATION: DE 1994-4435485 19941004

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: Patterson, Jr., Charles L.

LEGAL REPRESENTATIVE: Foley & Lardner

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

NUMBER OF DRAWINGS: 1 Drawing Figure(s); 1 Drawing Page(s)

LINE COUNT: 787

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB The invention relates to a method for isolation of highly pure von

Willebrand Factor in which recombinant von Willebrand Factor (rvWF) is chromatographically

purified by anion exchange chromatography on an anion exchanger of the quaternary amino type in a buffer solution comprising buffer substances and optionally salt.

The buffer solutions are preferably free of stabilizers, amino acids and other additives. According to this method, highly pure recombinant vWF can be obtained, which is free from blood plasma proteins, especially free from Factor VIII, and is physiologically active.

Further, the invention relates to a pharmaceutical preparation that contains rvWF, which is comprised of multimers with a high structural integrity.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 18 OF 72 USPATFULL on STN

ACCESSION NUMBER: 1999:27611 USPATFULL

TITLE: Method for isolation of highly pure von

Willebrand Factor

INVENTOR(S): Fischer, Bernhard, Vienna, Austria

> Mitterer, Artur, Orth/Donau, Austria Dorner, Friedrich, Vienna, Austria Schwarz, Hans-Peter, Vienna, Austria

Turecek, Peter, Vienna, Austria Eibl, Johann, Vienna, Austria

Falkner, Falko-Guenter, Orth/Donau, Austria

Schlokat, Uwe, Orth/Donau, Austria

Mundt, Wolfgang, Vienna, Austria Reiter, Manfred, Vienna, Austria

Den-Bouwmeester, Renate, Vienna, Austria

PATENT ASSIGNEE(S): Immuno Aktiengesellschaft, Vienna, Austria (non-U.S.

corporation)

NUMBER KIND DATE ______

US 5877152 19990302 US 1997-898131 19970722 (8) PATENT INFORMATION: APPLICATION INFO.: <--

RELATED APPLN. INFO.: Division of Ser. No. US 1996-653298, filed on 24 May

1996

NUMBER DATE

PRIORITY INFORMATION: DE 1994-4435485 19941004

WO 1995-EP3892 19951002

DOCUMENT TYPE: Utility

FILE SEGMENT: Granted
PRIMARY EXAMINER: Patterson, Jr., Charles L.

LEGAL REPRESENTATIVE: Foley & Lardner NUMBER OF CLAIMS: 9

EXEMPLARY CLAIM: NUMBER OF DRAWINGS:

1 Drawing Figure(s); 1 Drawing Page(s)

LINE COUNT: 767

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The invention relates to a method for isolation of highly pure von

Willebrand Factor in which recombinant von Willebrand Factor (rvWF) is chromatographically

purified by anion exchange chromatography on an anion exchanger of the quaternary amino type in a buffer solution comprising buffer substances and optionally salt.

The buffer solutions are preferably free of stabilizers, amino acids and other additives. According to this method, highly pure recombinant vWF can be obtained, which is free from blood plasma proteins, especially free from Factor VIII, and is physiologically active.

Further, the invention relates to a pharmaceutical preparation that contains rvWF, which is comprised of multimers with a high structural integrity

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 19 OF 72 USPATFULL on STN

ACCESSION NUMBER: 1999:15483 USPATFULL

Pharmaceutical preparation for treating blood TITLE:

coagulation disorders

INVENTOR(S): Turecek, Peter, Weidling, Austria

Schwarz, Hans-Peter, Vienna, Austria

<--

Eibl, Johann, Vienna, Austria

PATENT ASSIGNEE(S): Immuno Aktiengesellschaft, Vienna, Austria (non-U.S.

corporation)

NUMBER KIND DATE ______ PATENT INFORMATION: 19990202

US 5866122 US 1997-821763 19970320 (8) APPLICATION INFO.:

NUMBER DATE

______ AT 1996-518 19960320 AT 1996-1573 19960904 AT 1996-1673 19960920 PRIORITY INFORMATION:

AT 1996-1673

DOCUMENT TYPE: Utility

FILE SEGMENT: Granted
PRIMARY EXAMINER: Weddington, Kevin E. LEGAL REPRESENTATIVE: Foley & Lardner

NUMBER OF CLAIMS: 45 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 10 Drawing Figure(s); 8 Drawing Page(s)

LINE COUNT: 1609

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB There is disclosed a pharmaceutical preparation for treating blood coagulation disorders which comprises purified prothrombinase

factors, in particular purified prothrombin and optionally purified

factor Xa as active component.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L3 ANSWER 20 OF 72 SCISEARCH COPYRIGHT (c) 2005 The Thomson Corporation on

STN

ACCESSION NUMBER: 1999:239963 SCISEARCH

THE GENUINE ARTICLE: 176UU

TITLE: Preclinical evaluation of recombinant von

Willebrand factor in a canine model of

von Willebrand disease

AUTHOR: Schwarz H P (Reprint); Dorner F; Mitterer A; Mundt W;

Schlokat U; Pichler L; Turecek P L

CORPORATE SOURCE: Baxter Hyland Immuno, Ind Str 67, A-1220 Vienna, Austria

(Reprint); Baxter Hyland Immuno, A-1220 Vienna, Austria

COUNTRY OF AUTHOR: Austria

SOURCE: WIENER KLINISCHE WOCHENSCHRIFT, (12 MAR 1999)

Vol. 111, No. 5, pp. 181-191.

ISSN: 0043-5325.

PUBLISHER: SPRINGER-VERLAG WIEN, SACHSENPLATZ 4-6, PO BOX 89, A-1201

VIENNA, AUSTRIA.

DOCUMENT TYPE: Article; Journal

LANGUAGE:
REFERENCE COUNT:

English 41

ENTRY DATE: Entered STN: 1999

Last Updated on STN: 1999

ABSTRACT IS AVAILABLE IN THE ALL AND IALL FORMATS

AB Dutch Kooiker dogs with hereditary von Willebrand disease (vWD) have undetectable levels of von Willebrand factor (vWF),

resulting in spontaneous hemorrhage of mucosal surfaces similar to the clinical picture of vWD in humans. We used this canine model of vWD to study the in vivo effects of a new recombinant von Willebrand

factor (rvWF) preparation that contained all species of

 $vWF \ multimers \ compared \ with \ an \ rvWF \ fraction \ containing \ only \ low \ molecular \ weight \ multimers \ (LMW-rvWF) \ and \ with \ a \ plasma-derived \ factor \ VIII/vWF$

concentrate (pdvWF). Administration of rvWF in these

vWF-deficient dogs resulted in a vWF:Ag half-life of 21.6 hours in one dog and 22.1 hours in a second dog. Administration of pdvWF resulted in a half-life for vWF:Ag of 7.7 hours, and LMW-rvWF, 9 hours. The in vivo recovery of vWF:Ag after administration of rvWF was 59, 64 and 70% in three dogs, respectively; 33% after pdvWF, and 92% after LMW-rvWF. The in vivo recovery of ristocetin cofactor (RCoF) was 78, 110 and 120% for rvWF, and 25% for pdvWF. Both rvWF and pdvWF caused increases in factor VIII. Although no effect was seen on bleeding time at the dosages used, the rate of blood flow from cuticle wounds was reduced after a single bolus administration of rvWF. The rvWF was able to control a severe nose bleed in one dog.

L3 ANSWER 21 OF 72 USPATFULL on STN

ACCESSION NUMBER: 1998:162660 USPATFULL

TITLE: Method for isolation of highly pure von

Willebrand Factor

INVENTOR(S): Fischer, Bernhard, Vienna, Austria

Mitterer, Artur, Orth/Donau, Austria Dorner, Friedrich, Vienna, Austria Schwarz, Hans-Peter, Vienna, Austria

Turecek, Peter, Vienna, Austria Eibl, Johann, Vienna, Austria

Falkner, Falko-Guenter, Orth/Donau, Austria

Schlokat, Uwe, Orth/Donau, Austria

Mundt, Wolfgang, Vienna, Austria Reiter, Manfred, Vienna, Austria

Den-Bouwmeester, Renate, Vienna, Austria

<--

<--

Immuno Aktiengesellschaft, Vienna, Austria (non-U.S. PATENT ASSIGNEE(S):

corporation)

NUMBER KIND DATE -----

US 5854403 19981229 US 1996-653298 19960524 (8) PATENT INFORMATION:

APPLICATION INFO.:

NUMBER DATE -----

PRIORITY INFORMATION: DE 1994-4435485 19941004

DOCUMENT TYPE: Utility
FILE SEGMENT: Granted
PRIMARY EXAMINER: Patterson, Jr., Charles L.

LEGAL REPRESENTATIVE: Foley & Lardner

LEGAL REPRESENT...

NUMBER OF CLAIMS: 16

NUMBER OF DRAWINGS: 1 Drawing Figure(s); 1 Drawing Page(s)

LINE COUNT: 813

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The invention relates to a method for isolation of highly pure von

Willebrand Factor in which recombinant von Willebrand Factor (rvWF) is chromatographically

purified by anion exchange chromatography on an anion exchanger of the quaternary amino type in a buffer solution comprising buffer substances

and optionally salt. The buffer solutions are preferably free of stabilizers, amino acids and other additives. According to this method,

highly pure recombinant rvWF can be obtained, which is free from blood plasma proteins, especially free from Factor VIII, and is physiologically active. Further, the invention relates to a pharmaceutical preparation that contains rvWF, which comprises

mulitimers with a high structural integrity.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 22 OF 72 USPATFULL on STN

ACCESSION NUMBER: 1998:159734 USPATFULL

Process for producing a protein TITLE: INVENTOR(S): Adamson, Lars, Lidingo, Sweden

Walum, Erik, Akersberga, Sweden Dixelius, Johan, Uppsala, Sweden

Lie, Kristina Lima, Stockholm, Sweden

PATENT ASSIGNEE(S): Pharmacia & Upjohn AB, Stockholm, Sweden (non-U.S.

corporation)

NUMBER KIND DATE _____

PATENT INFORMATION: US 5851800 APPLICATION INFO.: US 1997-852783 19981222

19970507 (8)

NUMBER DATE _____

PRIORITY INFORMATION: SE 1996-1855 19960514
US 1996-18874P 19960529 (60)
DOCUMENT TYPE: Utility
FILE SEGMENT: Granted
PRIMARY EXAMINER: Weber, Jon P.

LEGAL REPRESENTATIVE: Dinsmore & Shohl LLP

19 NUMBER OF CLAIMS: 1 EXEMPLARY CLAIM: 719 LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

A process is presented for reducing the detrimental influence of certain proteases on recombinant human protein and polypeptide production in a cell culture, which comprises adding an inhibitor of metal-dependent

proteases or chymotrypsins to the cell culture medium. The cell culture medium for cultivating cells expressing and secreting a biologically active recombinant human polypeptide contains an inhibitor of metal-dependent proteases or chymotrypsins, or a combination thereof. Recombinant factor VIII which has been produced in a cell culture medium according to the present process is useful for the manufacture of a medicament for administration to a patient having the symptoms of hemophilia A and for treatment of hemophilia A by administration of a therapeutically effective amount of recombinant factor VIII.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

3 ANSWER 23 OF 72 USPATFULL on STN

ACCESSION NUMBER: 1998:107999 USPATFULL TITLE: Antiplasma animal model

TITLE: Antiplasma animal model INVENTOR(S): Eibl, Johann, Vienna, Austria

Turagel Poter Vlasterneuburg Weidl

Turecek, Peter, Klosterneuburg Weidling, Austria

Schwarz, Hans Peter, Vienna, Austria

PATENT ASSIGNEE(S): Immuno Aktiengesellschaft, Vienna, Austria (non-U.S.

corporation)

NUMBER KIND DATE

PATENT INFORMATION: US 5804159 19980908 <--

APPLICATION INFO.: US 1996-663031 19960607 (8)

NUMBER DATE

PRIORITY INFORMATION: AT 1995-987 19950609

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: Chambers, Jasemine C.

ASSISTANT EXAMINER: Hauda, Karen M. LEGAL REPRESENTATIVE: Foley & Lardner

NUMBER OF CLAIMS: 3 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 7 Drawing Figure(s); 7 Drawing Page(s)

LINE COUNT: 737

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

There is disclosed an anti-plasma antibody preparation for treatment of a mammal, which preparation is capable of directly or indirectly inhibiting and/or eliminating several blood factors, a method of producing such a preparation and a method of evaluating substances for treating clotting disorders by using said anti-plasma antibody preparation. There is further disclosed a method of determining the bleeding characteristics of a mammal.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L3 ANSWER 24 OF 72 USPATFULL on STN

ACCESSION NUMBER: 1998:33896 USPATFULL

TITLE: Composition comprising coagulation factor VIII

formulation, process for its preparation and

use of a surfactant as stabilizer Osterberg, Thomas, Stockholm, Sweden

INVENTOR(S): Osterberg, Thomas, Stockholm, Sweden Fatouros, Angelica, Stockholm, Sweden

PATENT ASSIGNEE(S): Pharmacia & Upjohn AB, Stockholm, Sweden (non-U.S.

corporation)

	NUMBER	KIND DATE	
PATENT INFORMATION:	บร 5733873	19980331	<
	WO 9407510	19940414	
APPLICATION INFO.:	US 1994-244266	19940520	(8)
•	WO 1993-SE793	19931001	
		19940520	PCT 371 date
		19940520	PCT 102(e) date

NUMBER DATE

PRIORITY INFORMATION: SE 1992-2878 19921002

SE 1993-1580 19930507 SE 1993-2006 19930611

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: Degen, Nancy

LEGAL REPRESENTATIVE: Pollock, Vande Sande & Priddy

NUMBER OF CLAIMS: EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 2 Drawing Figure(s); 1 Drawing Page(s)

LINE COUNT: 650

sugar alcohols.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

A composition comprising coaqulation factor VIII and a non-ionic surfactant such as block copolymers, e.g., polyoxamers or polyoxyethylene (20) sorbitan fatty acid esters, e.g., polysorbate 20 or polysorbate 80 as stabilizer is provided. The composition can also comprise sodium chloride, calcium chloride, L-histidine and/or sugars or

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 25 OF 72 SCISEARCH COPYRIGHT (c) 2005 The Thomson Corporation on 1.3

STN

AUTHOR:

1999:59508 SCISEARCH ACCESSION NUMBER:

THE GENUINE ARTICLE: 156NX

Evaluation of recombinant von Willebrand TITLE:

> factor in a canine model of von Willebrand disease Schwarz H P (Reprint); Dorner F; Mitterer A; Mundt W;

Schlokat U; Pichler L; Turecek P L

Immuno AG Wien, Industriestr 67, A-1220 Vienna, Austria CORPORATE SOURCE:

(Reprint); Baxter Healthcare Corp, Hyland Immuno Div,

Vienna, Austria

COUNTRY OF AUTHOR: Austria

HAEMOPHILIA, (1998) Vol. 4, Supp. [3], pp. 53-62 SOURCE:

ISSN: 1351-8216.

BLACKWELL SCIENCE LTD, P O BOX 88, OSNEY MEAD, OXFORD OX2 PUBLISHER:

> ONE, OXON, ENGLAND. Article; Journal

LANGUAGE: English

DOCUMENT TYPE:

REFERENCE COUNT: 35

Entered STN: 1999 ENTRY DATE:

Last Updated on STN: 1999

ABSTRACT IS AVAILABLE IN THE ALL AND IALL FORMATS

Dutch Kooiker dogs with hereditary von Willebrand disease have undetectable levels of von Willebrand factor (vWF), resulting in spontaneous haemorrhage of mucosal surfaces similar to the clinical picture of von Willebrand disease in humans. We used this canine model of von Willebrand disease to study the in vivo effects of a new recombinant von Willebrand factor (rvWF)

preparation that contained all species of vWF multimers compared with a rvWF fraction containing only low molecular weight multimers (LMW-rvWF) and with a plasma-derived factor VIII/vWF concentrate (pdvWF). Administration of rvWF in these vWF-deficient dogs resulted in a vWF: Ag half-life of 21.6 h in one dog and 22.1 h in a second dog. Administration of pdvWF resulted in a half-life for vWF:Ag of 7.7 h, and LMW-rvWF, 9 h. The in vivo recovery of vWF:Ag after administration of rvWF was 59%, 64% and 70% in three dogs, respectively; 33% after pdvWF, and 92% after LMW-rvWF. The in vivo recovery of ristocetin cofactor (RCoF) was 78%, 110% and 120% for rvWF, and 25% for pdvWF. Both rvWF and pdvWF caused increases in FVIII. Although no effect was seen on bleeding time at the dosages used, the rate of blood flow from cuticle wounds was reduced after a single bolus administration of rvWF. The rvWF was able to control a severe nose bleed in one dog.

AΒ

AN 02807564 IFIPAT;IFIUDB;IFICDB

TITLE: FACTOR VIII BINDING DOMAIN OF VON WILLEBRAND

FACTOR

INVENTOR(S): Foster, Paul A, San Diego, CA

Fulcher, Carol A, La Jolla, CA

Zimmerman, Theodore S, La Jolla, CA

PATENT ASSIGNEE(S): The Scripps Research Institute, La Jolla, CA

PRIMARY EXAMINER: Jacobson, Dian C

NUMBER PK DATE

PATENT INFORMATION: US 5597711 A 19970128 (CITED IN 001 LATER PATENTS)

TO THE ALONG ALONG

APPLICATION INFORMATION: US 1995-410574 19950324

EXPIRATION DATE: 28 Jan 2014

GRANTED PATENT NO. DATE APPLN. NUMBER OR STATUS _____ _____ 19870501 5043429 US 1987-45032 CONTINUATION OF: US 1993-125559 19930923 ABANDONED CONTINUATION OF: US 1991-725560 19910703 5260274 DIVISION OF: US 5597711 FAMILY INFORMATION: 19970128 US 5043429 US 5260274 Utility DOCUMENT TYPE: FILE SEGMENT: CHEMICAL

NUMBER OF CLAIMS: 6

AB Peptides which inhibit the binding of yon Willebrand
Factor to Factor VIII. Monoclonal antibodies capable of
specifically binding to the region of von Willebrand

GRANTED

Factor containing the Factor VIII binding domain. Improved

methods of preparing Factor VIII.

CLMN 6

L3 ANSWER 27 OF 72 USPATFULL on STN

ACCESSION NUMBER: 97:99199 USPATFULL

TITLE: Retroviral delivery of full length factor VIII INVENTOR(S): Bodner, Mordechai, San Diego, CA, United States

Bodner, Mordechai, San Diego, CA, United States De Polo, Nicholas J., Solana Beach, CA, United States

Chang, Stephen, Poway, CA, United States

Hsu, David Chi-Tang, San Diego, CA, United States Respess, James G., San Diego, CA, United States

<--

PATENT ASSIGNEE(S): Chiron Viagene, Inc., United States (U.S. corporation)

NUMBER KIND DATE

PATENT INFORMATION: US 5681746 19971028 APPLICATION INFO.: US 1994-366851 19941230 (8)

DOCUMENT TYPE: Utility
FILE SEGMENT: Granted

FILE SEGMENT: Granted
PRIMARY EXAMINER: Crouch, Deborah
ASSISTANT EXAMINER: Schmuck, Jill

LEGAL REPRESENTATIVE: Kruse, Norman J., Blackburn, Robert P.

NUMBER OF CLAIMS: 14 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 5 Drawing Figure(s); 5 Drawing Page(s)

LINE COUNT: 3229

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB Retroviral vectors for directing expression of full length factor VIII in transduced host cells, plasmids encoding the same, and host cells transformed, transfected, or transduced therewith are disclosed. Also disclosed are retroviral particles comprising such retrovital vectors, as are methods for making such particles in suitable packaging cells. Retroviral particles so produced may be amphotropic, ecotropic, polytropic, or xenotropic; alternatively, they may comprise chimeric or

hybrid envelope proteins to alter host range. Also described are retrovital particles comprising retroviral vectors for directing full length factor VIII expression which are complement resistant. Pharmaceutical compositions comprising retrovital particles of the invention are also disclosed, as are methods of treating mammals, particularly humans, afflicted with hemophilia.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L3 ANSWER 28 OF 72 USPATFULL on STN

ACCESSION NUMBER: ~97:75993 USPATFULL

TITLE: Recombinant human factor VIII derivatives INVENTOR(S): Almstedt, Annelie B., Stockholm, Sweden

Gray (Hellstrom), Eva Maria, Stockholm, Sweden

Lind, Peter, Upsala, Sweden

Ljung, Catherine, Vallingby, Sweden Sandberg, Helena Inga, Bromma, Sweden

Spira, Jack, Solna, Sweden

Sydow-Backman, Mona, Saltsjobaden, Sweden

Wiman, Helena, Stockholm, Sweden

PATENT ASSIGNEE(S): Kabi Pharmacia AB, Upsala, Sweden (non-U.S.

corporation)

NUMBER KIND DATE

PATENT INFORMATION: US 5661008 19970826 <--

APPLICATION INFO.: US 1995-462917 19950605 (8)

RELATED APPLN. INFO.: Continuation of Ser. No. US 1992-934495, filed on 17

Dec 1992, now abandoned

NUMBER DATE

PRIORITY INFORMATION: SE 1991-799 19910315

DOCUMENT TYPE: Utility
FILE SEGMENT: Granted

PRIMARY EXAMINER: Jacobson, Dian C.

LEGAL REPRESENTATIVE: Burns, Doane, Swecker & Mathis, L.L.P.

NUMBER OF CLAIMS: 13 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 8 Drawing Figure(s); 8 Drawing Page(s)

LINE COUNT: 833

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB A DNA sequence coding for a biologically active recombinant human factor VIII derivative, comprising a first DNA segment coding for the amino acids 1 through 740 of human factor VIII and a second DNA segment coding for the amino acids 1649 through 2332 of human factor VIII, said segments being interconnected by a linker DNA segment coding for a linker peptide of at least 3 amino acid residues and up to about 10 amino acid residues which are selected from lysine and arginine; recombinant expression vector comprising such DNA sequence; host cells of animal origin transformed with such recombinant expression vector; a process for the manufacture of recombinant human factor VIII derivative; and human factor VIII derivative containing the heavy chain and the light chain linked by metal ion bond.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L3 ANSWER 29 OF 72 USPATFULL on STN

PATENT ASSIGNEE(S):

ACCESSION NUMBER: 97:73722 USPATFULL

TITLE: Anion exchange process for the purification of Factor

VIII

INVENTOR(S): Bhattacharya, Prabir, Walnut, CA, United States

Motokubota, Toshiharu, Arcadia, CA, United States

Fedalizo, Norman M., Rowland Heights, CA, United States Alpha Therapeutic Corporation, Los Angeles, CA, United

States (U.S. corporation)

NUMBER KIND DATE

PATENT INFORMATION: US 5659017 19970819 <--

APPLICATION INFO.: US 1995-554724 19951107 (8)

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: Schain, Howard E.

LEGAL REPRESENTATIVE: Christie, Parker & Hale, LLP

NUMBER OF CLAIMS: 23 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 11 Drawing Figure(s); 8 Drawing Page(s)

LINE COUNT: 866

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB A new method for purifying Factor VIII complex from an impure protein fraction, usually cryoprecipitate, is disclosed. The cryoprecipitate is dissolved in a heparin solution. Then Factor VIII complex is initially purified by polyethylene glycol precipitation. The Factor VIII containing supernatant collected after the precipitation is loaded into an anion exchange column that has a quaternary amino ethyl group. The Factor VIII complex is then eluted from the column with a buffer comprising from about 0.14M to about 0.20M CaCl.sub.2. The final step in the purification is to precipitate the Factor VIII complex in the presence of glycine and sodium chloride. The precipitated Factor VIII complex is then reconstituted and stabilized. The reconstituted Factor VIII complex can then be lyophilized and dry heated to obtain a final Factor VIII product.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L3 ANSWER 30 OF 72 DRUGU COPYRIGHT 2005 THE THOMSON CORP on STN

ACCESSION NUMBER: ,1997-35135 DRUGU T

TITLE: Successful cardiac surgery in a young girl with type 2a von

Willebrand disease using continuous infusion of Hemate-P.

AUTHOR: Naqvi A; Endres Brooks J; Montgomery R R; Stain A M; Sparling

C; Klaassen R; Blanchette V S

LOCATION: Toronto, Ont., Can.; Milwaukee, Wis., USA SOURCE: Thromb.Haemostasis (Suppl., 514, 1997) 1 Ref.

CODEN: THHADQ ISSN: 0340-6245

AVAIL. OF DOC.: Division of Hematology, the Hospital for Sick Children,

Toronto, Ontario, Canada.

LANGUAGE: English
DOCUMENT TYPE: Journal
FIELD AVAIL: AB; LA; CT
FILE SEGMENT: Literature
AN 1997-35135 DRUGU T

AB Here the Authors report the successful use of continuous infusion therapy using a factor VIII/von Willebrand factor

concentrate (Hemate-P) to provide hemostatic cover in a 6-yr-old girl with type 2A von Willebrand disease (vWd) admitted for elective closure of a perimembranous ventriculoseptal defect using temporary cardiopulmonary bypass and moderate hypothermia. The patient also received desmopressin (DDAVP) during the preoperative period. The patient experienced no untoward bleeding in the operative or post-operative periods and no infectious complications were noted. (conference abstract).

ABEX Baseline coagulation results were as follows: platelet count 345 x 10 power 9/1, bleeding time 17 min (normal less than 7 min), FVIII:C 1.47 U/ml; vWf:Ag 0.84 U/ml, and vWf R:Co 0.42 U/ml. Preoperatively the patient received 74 U/kg of Hemate-P and 0.3 ug/kg of DDAVP by i.v. infusion with correction of the bleeding time and haemostatic parameters (post infusion bleeding time 6 min; FVIII:C 4.63 U/ml; vWf:Ag 2.25 U/ml; and vWf R:Co 2.06 U/ml). Postoperatively the patient was maintained on a continuous infusion of Hemate-P for 7 days (initial rate 2.7 U/kg/hr, concentration 25 U/ml, decreasing to 0.4 U/kg/hr, concentration 12.5 U/ml). Hemate-P was reconstituted as per manufacturers instruction, dispensed in a 60 ml syringe and administered through a high-precision infusion pump. The one exception was the last concentrate preparation; this material was diluted 1:1 with normal saline (final concentration 6.5 U/ml). Coagulation testing on samples of

reconstituted Hemate-P (multiple lots) demonstrated near normal distribution of vWf multimers for up to 40 hr post reconstitution. Following infusion of the Hemate-P, the patient's vWf multimeric pattern nearly normalized with reconstitution of the high molecular weight

multimeric forms. Sustained levels of both von-Willebrand

factor and factor VIII:C were kept

at hemostatic levels or above through the continuous infusion of Hemate-P in this patient with type 2A von Willebrand disease. The ratio of vWf R:Co and vWf:Ag were similar during the treatment period. (PH)

ANSWER 31 OF 72 USPATFULL on STN

ACCESSION NUMBER: 96:94557 USPATFULL

Stabilized factor VIII preparations TITLE:

INVENTOR(S): Freudenberg, Wilfried, C olbe-Sch onstadt, Germany,

Federal Republic of

PATENT ASSIGNEE(S): Behringwerke Aktiengesellschaft, Marburg, Germany,

Federal Republic of (non-U.S. corporation)

NUMBER KIND DATE -----

US 5565427 · PATENT INFORMATION: US 5565427 19961015 APPLICATION INFO.: US 1994-235241 19940429 (8)

RELATED APPLN. INFO.: Continuation of Ser. No. US 1993-82911, filed on 29 Jun 1993, now abandoned which is a continuation of Ser. No.

US 1992-864610, filed on 7 Apr 1992, now abandoned

NUMBER DATE ______

PRIORITY INFORMATION: DE 1991-4111393 19910409

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: PRIMARY EXAMINER: Weimar, Elizabeth C. ASSISTANT EXAMINER: Touzeau, P. Lynn

LEGAL REPRESENTATIVE: Finnegan, Henderson, Farabow, Garrett & Dunner, L.L.P.

LEGAL REPRESENTATIVE.

NUMBER OF CLAIMS: 13 238 LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The invention relates to stabilized solutions with F VIII coagulation

activity, to a process for the preparation thereof and to the

use thereof.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 32 OF 72 USPATFULL on STN L3

ACCESSION NUMBER: 96:5889 USPATFULL

Antihemophilic factor stabilization TITLE:

Johnson, Alan J., 127 W. 12th St., New York, NY, United INVENTOR(S):

States 10011

Fulton, Anne J., 515 Avon Dr., East Windsor, NJ, United

States 08520

NUMBER KIND DATE ______

US 5484890 PATENT INFORMATION: 19960116

APPLICATION INFO.: US 1993-138481 19931015 (8)

DISCLAIMER DATE: 20110111

RELATED APPLN. INFO.: Continuation of Ser. No. US 1991-790390, filed on 12

Nov 1991, now patented, Pat. No. US 5278289

DOCUMENT TYPE: Utility

FILE SEGMENT: Granted PRIMARY EXAMINER: Russel, Jeffrey E. LEGAL REPRESENTATIVE: Darby & Darby

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

NUMBER OF DRAWINGS: 1 Drawing Figure(s); 1 Drawing Page(s)

LINE COUNT: 927

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

A method of recovering a purified and stabilized protein having AB antihemophilic factor activity (Factor VIII), from a biological sample containing Factor VIII, at least one destabilizing protease impurity, i.e. thrombin, and optionally one or more proprotease impurity, is provided. The biological sample is contacted with a protease removing and/or inhibiting agent, thereby inhibiting and/or removing the destabilizing protease impurity. The method provides increased yield and resolution of Factor VIII. Also provided are purified and stabilized Factor VIII non-lyophilized, liquid in compositions.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 33 OF 72 USPATFULL on STN

95:105944 USPATFULL ACCESSION NUMBER:

Ultrapurification process for factor VIII TITLE:

Neslund, Gerard G., Diamond Bar, CA, United States INVENTOR(S):

Liu, Shu-Len, Cerritos, CA, United States

Griffith, Michael J., Claremont, CA, United States

Baxter International Inc., Deerfield, IL, United States PATENT ASSIGNEE(S):

(U.S. corporation)

NUMBER KIND DATE

 US
 5470954
 19951128

 US
 1993-140695
 19931021

 PATENT INFORMATION:

APPLICATION INFO.: 19931021 (8)

Continuation of Ser. No. US 1992-887387, filed on 21 RELATED APPLN. INFO.:

May 1992, now abandoned which is a continuation of Ser. No. US 1988-167902, filed on 28 Mar 1988, now abandoned

which is a continuation-in-part of Ser. No. US 1987-32800, filed on 31 Mar 1987, now abandoned

DOCUMENT TYPE: Granted FILE SEGMENT:

PRIMARY EXAMINER: Sayala, Chhaya D. LEGAL REPRESENTATIVE: Condino, Debra D.

NUMBER OF CLAIMS: 39 EXEMPLARY CLAIM: 1 1233 LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT. A process for purifying Factor VIII:C

comprising contacting an immobilized antibody specifically binding a

Factor VIII: C with Factor VIII: C, desorbing Factor VIII:

c from the antibody which had adsorbed it, eluting

Factor VIII:C from the presence of the antibody, passing the eluted Factor VIII:C

through an affinity region capable of binding the Factor

VIII:C, binding the Factor VIII:

C in the affinity region and passing contaminants through said

region, and eluting the purified Factor VIII:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 34 OF 72 USPATFULL on STN

ACCESSION NUMBER: 95:25015 USPATFULL

Solubilization and stabilization of factor VIII complex TITLE:

INVENTOR(S): Bhattacharya, Prabir, Walnut, CA, United States Motokubota, Toshiharu, Arcadia, CA, United States

PATENT ASSIGNEE(S): Alpha Therapeutic Corporation, Los Angeles, CA, United

States (U.S. corporation)

KIND NUMBER DATE ------

US 5399670 19950321 US 1993-54903 19930429 (8) PATENT INFORMATION:

APPLICATION INFO.:

Continuation-in-part of Ser. No. US 1992-876190, filed RELATED APPLN. INFO.:

on 30 Apr 1992, now abandoned

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: Schain, Howard E. ASSISTANT EXAMINER: Touzeau, Lynn

Christie, Parker & Hale LEGAL REPRESENTATIVE:

NUMBER OF CLAIMS: EXEMPLARY CLAIM: LINE COUNT: 524

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

A process for facilitating the reconstitution of lyophilized Factor VIII complex compositions, and compositions of Factor VIII complex, which are readily reconstituted. The process of the present invention comprises providing a purified Factor VIII complex preparations; adding a stabilization agent comprising arginine; lyophilizing the stabilization agent-Factor VIII complex solutions; and reconstituting the lyophilized stabilization agent-Factor VIII complex by contacting it with solvent for less than one minute.

'CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 35 OF 72 USPATFULL on STN.

ACCESSION NUMBER: 94:91035 USPATFULL

TITLE: Gel filtration of factor VIII

Brockway, William J., Oakland, CA, United States INVENTOR(S):

Seng, Richard L., Guerneville, CA, United States

Miles Inc., Berkeley, CA, United States (U.S. PATENT ASSIGNEE(S):

corporation)

NUMBER KIND DATE _____

US 5356878 PATENT INFORMATION: 19941018

US 1993-852 19930104 (8) APPLICATION INFO.:

Division of Ser. No. US 1990-587815, filed on 24 Sep RELATED APPLN. INFO.: 1990, now patented, Pat. No. US 5177591 which is a

continuation of Ser. No. US 1987-135966, filed on 21

Dec 1987, now abandoned

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: Schain, Howard E. ASSISTANT EXAMINER: Touzeau, P. Lynn

LEGAL REPRESENTATIVE: Giblin, James A., Bradley, Bertram

NUMBER OF CLAIMS: EXEMPLARY CLAIM: LINE COUNT: 693

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Highly purified antihemophilic factor is produced by a process comprising a PEG precipitation step, a gel filtration step and a virus inactivation step. Al (OH).sub.3 adsorption and PEG precipitation carried out at room temperature allow processing to proceed directly to a gel filtration step.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 36 OF 72 USPATFULL on STN

ACCESSION NUMBER: 94:15877 USPATFULL

Factor viii purification process TITLE:

INVENTOR(S): Bhattacharva, Prabir, Walnut, CA, United States

Motokubota, Toshiharu, Arcadia, CA, United States

<--

Alpha Therapeutic Corporation, Los Angeles, CA, United PATENT ASSIGNEE(S):

States (U.S. corporation)

NUMBER KIND DATE -----

US 5288853 19940222 PATENT INFORMATION: 19920430 (7) US 1992-876410 APPLICATION INFO.:

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

Schain, Howard R. PRIMARY EXAMINER: Touzeau, Lynn ASSISTANT EXAMINER:

LEGAL REPRESENTATIVE: Christie, Parker & Hale

NUMBER OF CLAIMS: 21 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 2 Drawing Figure(s); 2 Drawing Page(s)

LINE COUNT: 777

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

There is provided in accordance with the practice of this invention a process for separating Factor VIII complex from an impure protein fraction containing Factor VIII complex. An aqueous solution of the impure protein fraction containing Factor VIII complex is applied to a heparin-coupled chromatographic medium, to bind the Factor VIII complex to the medium. The Factor VIII is then recovered from the heparin-coupled chromatographic medium by elution with an aqueous solution comprising CaCl.sub.2 and histidine. The recovered Factor VIII is further purified by precipitation with a solution comprising glycine and NaCl, and washing the resultant precipitate with a solution comprising histidine, glycine, and NaCl to provide a Factor VIII complex solution with a specific activity of about 70 to about 150 units/mg.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L3 ANSWER 37 OF 72 USPATFULL on STN ACCESSION NUMBER: 94:3911 USPATFULL

TITLE: Antihemophilic factor stabilization

INVENTOR(S): Johnson, Alan J., 127 W. 12th St., New York, NY, United

States 10011

Fulton, Anne J., 515 Avon Dr., East Windsor, NJ, United

States 08520

NUMBER KIND DATE

PATENT INFORMATION: US 5278289 19940111 <--

APPLICATION INFO.: US 1991-790390 19911112 (7)

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: Russel, Jeffrey E. LEGAL REPRESENTATIVE: Darby & Darby

NUMBER OF CLAIMS: 2 EXEMPLARY CLAIM: 1

INVENTOR(S):

NUMBER OF DRAWINGS: 1 Drawing Figure(s); 1 Drawing Page(s)

LINE COUNT: 756

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB A method of recovering a purified and stabilized protein having antihemophilic factor activity (Factor VIII), from a biological sample containing Factor VIII, at least one destabilizing protease impurity, i.e. thrombin, and optionally one or more proprotease impurity, is provided. The biological sample is contacted with a protease removing and/or inhibiting agent, thereby inhibiting and/or removing the destabilizing protease impurity. The method provides increased yield and resolution of Factor VIII. Also provided are purified and stabilized Factor VIII non-lyophilized, liquid in compositions.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L3 ANSWER 38 OF 72 USPATFULL on STN

ACCESSION NUMBER: 93:93445 USPATFULL

TITLE: Process for the purification of factor VIII and factor

VIII obtained by said process Arrighi, Silvana, Rieti, Italy

Borri, Maria G., Siena, Italy Ceccarini, Costante, Castelnuovo Berardenga, Italy

PATENT ASSIGNEE(S): SCLAVO S.p.A., Siena, Italy (non-U.S. corporation)

NUMBER KIND DATE

PATENT INFORMATION: US 5259951 19931109 <-APPLICATION INFO.: US 1991-713071 19910611 (7)

NUMBER DATE -----

PRIORITY INFORMATION: IT 1990-20610 19900612

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: Dawson, Robert A.
ASSISTANT EXAMINER: Kim, Sun Uk PRIMARY EXAMINER:

LEGAL REPRESENTATIVE: Birch, Stewart, Kolasch & Birch

NUMBER OF CLAIMS: 21 EXEMPLARY CLAIM: 1 307 LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

A method for the purification of Factor VIII from human plasma is described, wherein a solution comprising Factor VIII is purified by using ion exchange chromatographic columns. Factor VIII obtained by said

method is also described.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L3 ANSWER 39 OF 72 USPATFULL on STN

ACCESSION NUMBER: 93:76632 USPATFULL

TITLE: Method for isolating factors VIII from plasma by gel

filtration chromatography under group separation

conditions

Kaersgaard, Per, Vedbaek, Denmark INVENTOR(S):

PATENT ASSIGNEE(S): Novo Nordisk A/S, Bagsvaerd, Denmark (non-U.S.

corporation)

NUMBER KIND DATE ______

PATENT INFORMATION: US 5245014 19930914 APPLICATION INFO.: US 1990-610480 19901107 <--

19901107 (7)

NUMBER DATE _____

PRIORITY INFORMATION: DK 1989-5621 19891109

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: Wax, Robert A.
ASSISTANT EXAMINER: Ekstrom, Richard C.

LEGAL REPRESENTATIVE: Zelson, Steve T., Lambiris, Elias J.

NUMBER OF CLAIMS: 6 1 EXEMPLARY CLAIM:

NUMBER OF DRAWINGS: 2 Drawing Figure(s); 2 Drawing Page(s)

598 LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

A method for isolating Factor VIII from other proteins dissolved in blood plasma is disclosed, wherein plasma is subjected to gel filtration under group separation conditions giving a fraction containing Factor VIII in very high yield and almost free of other proteins.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 40 OF 72 USPATFULL on STN

ACCESSION NUMBER: 93:27199 USPATFULL

TITLE: Method for purifying factor VIII:

> C, von Willebrand factor and complexes thereof .

Kumar, Anur A., Seattle, WA, United States INVENTOR(S):

Hagen, Frederick S., Seattle, WA, United States Sledziewski, Andrzej Z., Seattle, WA, United States

ZymoGenetics, Inc., Seattle, WA, United States (U.S. PATENT ASSIGNEE(S):

corporation)

KIND DATE NUMBER ______

US 5200510 19930406 PATENT INFORMATION: <--US 1988-162877 19880302 (7) APPLICATION INFO.:

Continuation-in-part of Ser. No. US 1987-62896, filed RELATED APPLN. INFO.:

on 16 Jun 1987, now abandoned

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: Wax, Robert A. Furman, Keith C. ASSISTANT EXAMINER: LEGAL REPRESENTATIVE: Seed and Berry

NUMBER OF CLAIMS: 21 EXEMPLARY CLAIM:

NUMBER OF DRAWINGS: 2 Drawing Figure(s); 2 Drawing Page(s)

LINE COUNT: 626

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Methods for purifying factor VIII:C, von .

Willebrand factor (vWF) or complexes thereof from

heterogeneous biological fluids are disclosed. The methods utilize a

binding peptide, specific to either factor VIII:

c or vWF, bound to an insoluble matrix. Peptides suitable for

use within the methods are also disclosed.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 41 OF 72 USPATFULL on STN ACCESSION NUMBER: 93:5245 USPATFULL

Method for the treatment of bleeding disorders TITLE:

Hedner, Ulla K. E., Bagangsvagen 29, SE-21620 Malmo, INVENTOR(S):

Sweden

NUMBER KIND DATE · -----

US 5180583 PATENT INFORMATION: 19930119

US 1991-666423 19910308 (7) APPLICATION INFO.:

Continuation of Ser. No. US 1989-368967, filed on 19 RELATED APPLN. INFO.: Jun 1989, now abandoned which is a continuation of Ser.

No. US 1986-933408, filed on 20 Nov 1986, now abandoned

NUMBER DATE

DK 1985-5446 19851126 DK 1986-459285 19860926 PRIORITY INFORMATION:

DOCUMENT TYPE: FILE SEGMENT: Utility

FILE SEGMENT: Granted
PRIMARY EXAMINER: Stone, Jacqueline

Zelson, Steve T., Lambiris, Elias J. LEGAL REPRESENTATIVE:

NUMBER OF CLAIMS: EXEMPLARY CLAIM: 1 LINE COUNT: 463

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

A method for treating patients suffering from bleeding disorders not caused by clotting factor defects or clotting factor inhibitors, as well as a novel composition for use in treating bleeding disorders as disclosed. The method includes administering to a patient a composition comprising an effective haemostatic amount of factor VIIa, and is particularly effective in treating patients suffering from thrombocytopenia and von Willebrand's disease, as well as other platelet disorders. A composition suitable for use in treating such bleeding disorders comprises purified factor VIIa in a concentration of at least $25 \mu g/ ml.$

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 42 OF 72 USPATFULL on STN 93:1491 USPATFULL ACCESSION NUMBER:

GET filtration of factor VIII TITLE:

Brockway, William J., Oakland, CA, United States INVENTOR(S):

Seng, Richard L., Guerneville, CA, United States

Miles, Inc., Elkhart, IN, United States (U.S. PATENT ASSIGNEE(S):

corporation)

NUMBER KIND DATE _______

.19930105 19900924 (7) PATENT INFORMATION: US 5177191 US 1990-587815 <--

APPLICATION INFO.:

Continuation of Ser. No. US 1987-135966, filed on 21 RELATED APPLN. INFO.:

Dec 1987, now abandoned

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: Cashion, Jr., Merrell C.

ASSISTANT EXAMINER: Touzeau, P. Lynn

Aston, David J., Bradley, Bertram, Giblin, James A. LEGAL REPRESENTATIVE:

NUMBER OF CLAIMS: EXEMPLARY CLAIM: 1 683 LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Highly purified antihemophilic factor can be produced from a cryoprecipitate by a process comprising a PEG precipitation step, a viral inactivation step and a gel filtration step, all steps being carried out at room temperature.

ANSWER 43 OF 72 JICST-EPlus COPYRIGHT 2005 JST on STN

ACCESSION NUMBER: 930674621 JICST-EPlus

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Effect of Substrate and Diluent for Factor VIII Activity in TITLE:

> Highly Purified Factor VIII Concentrate and Patient Plasma Infused Factor VIII Concentrate.

TAKAHASHI ISAO; FURUTA MOTOMU; MIZUNO SHIN'ICHI; KAMIYA AUTHOR:

TADASHI

TAKAMATSU JUNKI

Aichi Red Cross Blood Center CORPORATE SOURCE:

Nagoya Univ., School of Medicine

Rinsho Byori (Japanese Journal of Clinical Pathology), SOURCE:

(1993) vol. 41, no. 7, pp. 825-830. Journal Code: Z0687A

(Fig. 1, Tbl. 3, Ref. 8) CODEN: RBYOAI; ISSN: 0047-1860

PUB. COUNTRY: Japan

DOCUMENT TYPE: Journal; Article

LANGUAGE: Japanese STATUS: New

We studied the difference between congenital factor VIII deficient plasma and factor VIII immunodepleted plasma on the effect of predilution for factor VIII activity determined by the one-stage assay. When standard

curves of one-stage assay for factor VIII: C

by GEORGE KING factor VIII deficient plasma (frozen at -80.DEG.C.), BEHRING factor VIII deficient plasma (lyophilized), DADE factor VIII

depleted plasma (lyophilized, von Willebrand factor

antigen 0.2 U/ml) and DADE factor VIII depleted plasma (lyophilized, von

Willebrand factor antigen 1.0 U/ml) were compared, the

difference between the clotting times for 100% and 6.25% of activity in each reagent were 39.5, 29.5, 25.0, 23.0 seconds respectively. Potency

values in concentrates without albumin or von Willebrand

factor showed a discrepancy betweenpredilution in Owren-Koller buffer and predilution in factor VIII deficient plasma. Potencies of those products prediluted in Owren-Koller buffer were 40-60% lower than potencies prediluted in factor VIII deficient plasma. These results showed substrate and prediluent must be chosen carefully for the accurate assay of factor VIII activity in vitro for the highly purified factor VIII

concentrates. (author abst.)

ANSWER 44 OF 72 USPATFULL on STN

ACCESSION NUMBER: 92:63798 USPATFULL

TITLE: Method for evaluating immunogenicity

Esmon, Pamela C., Richmond, CA, United States INVENTOR(S):

Fournel, Michael A., Castro Valley, CA, United States Miles Inc., Elkhart, IN, United States (U.S.

PATENT ASSIGNEE(S):

corporation)

NUMBER KIND DATE ______

PATENT INFORMATION:
APPLICATION INFO.: US 5135874 US 5135874 19920804 US 1990-493659 19900315 (7) <--

RELATED APPLN. INFO.: Continuation-in-part of Ser. No. US 1988-202177, filed

on 1 Jun 1988

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: Kepplinger, Esther L. ASSISTANT EXAMINER: Bidwell, Carol E.

Aston, David J., Enayati, Elizabeth F. LEGAL REPRESENTATIVE:

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

NUMBER OF DRAWINGS: 9 Drawing Figure(s); 8 Drawing Page(s)

LINE COUNT: 897

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

A method for evaluating the potential immunogenicity of a protein derived from recombinant DNA technology. The method involves injecting an animal with the recombinant protein and then isolating antiserum from the animal. The antiserum is depleted of antibodies to a reference protein, i.e., a plasma derived protein, by adsorbing the antiserum against the reference protein. The adsorbed antiserum is then blotted against the recombinant protein, to see if any antibodies were produced which recognize the recombinant protein, but did not recognize the plasma-derived protein during adsorption.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 45 OF 72 USPATFULL on STN

92:36292 USPATFULL ACCESSION NUMBER:

Factor VIII complex purification using heparin affinity TITLE:

chromatography

Kosow, David P., Monrovia, CA, United States INVENTOR(S):

Bhattacharya, Prabir, Walnut, CA, United States Sternburg, Charles F., Norco, CA, United States

Alpha Therapeutic Corporation, Los Angeles, CA, United PATENT ASSIGNEE(S):

States (U.S. corporation)

NUMBER KIND DATE ______

US 5110907 US 1989-388254 19920505 <--

PATENT INFORMATION: APPLICATION INFO.: 19890801 (7)

DOCUMENT TYPE: Utility Granted FILE SEGMENT:

PRIMARY EXAMINER: PRIMARY EXAMINER: Wax, Robert A.
ASSISTANT EXAMINER: Ekstrom, Richard C. LEGAL REPRESENTATIVE: Christie, Parker & Hale

NUMBER OF CLAIMS: 15 EXEMPLARY CLAIM: LINE COUNT: 842

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

There is provided in accordance with the practice of this invention, a process for separating Factor VIII complex from an impure protein fraction containing Factor VIII complex. An aqueous solution of the impure protein fraction containing Factor VIII complex is applied to a heparin coupled chromatographic medium to bind the Factor VIII complex to the medium. The Factor VIII is then recovered from the heparin coupled chromatographic medium by elution with an aqueous CaCl.sub.2 solution.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 46 OF 72 USPATFULL on STN

<u>92:1</u>4989 USPATFULL ACCESSION NUMBER:

Agent for the therapy of factor VIII-resistant TITLE:

hemophilia A, and a process for the preparation

thereof

Heimburger, Norbert, Marburg, Germany, Federal Republic INVENTOR(S):

οf

Wenz, Karlheinz, Weimar, Germany, Federal Republic of Wormsbacher, Wilfried, Kirchhain, Germany, Federal

Republic of

Behringwerke Aktiengesellschaft, Marburg/Lahn, Germany, PATENT ASSIGNEE(S):

Federal Republic of (non-U.S. corporation)

KIND DATE -----

US 5091363 19920225 US 1988-230717 19880810 (7) PATENT INFORMATION:

APPLICATION INFO.:

RELATED APPLN. INFO.: Continuation of Ser. No. US 1987-76600, filed on 22 Jul

1987, now abandoned

NUMBER DATE ______

PRIORITY INFORMATION:

DE 1986-3625090 19860724

DOCUMENT TYPE: Utility Granted FILE SEGMENT: Granted Rosen, Sam PRIMARY EXAMINER:

LEGAL REPRESENTATIVE: Finnegan, Henderson, Farabow, Garrett, and Dunner

NUMBER OF CLAIMS: 6 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 1 Drawing Figure(s); 1 Drawing Page(s)

LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

An agent for the therapy of hemophilia A which is resistant to treatment with factor VIII is described, and is obtainable by maintaining a mixture of factor VIII, antithrombin III, a phospholipid and calcium ions in an aqueous solution at a temperature of from 1° to 45° C. for at least one minute, adding factor IX, and maintaining the solution at a temperature offrom 1° to 45° C. until addition of a sample of this solution to an inhibitor plasma results in a partial thromboplastin time (PTT) of 15 to 30 seconds, where appropriate adding a polyol and, where appropriate, an amino acid, and,

where appropriate, drying the solution.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 47 OF 72 USPATFULL on STN

91:44553 USPATFULL ACCESSION NUMBER:

TITLE: Process for the preparation of factor

VIII:C-deficient plasma, and a

deficient plasma obtained in this way

INVENTOR(S): Becker, Udo, Munchen, Germany, Federal Republic of

Heimburger, Norbert, Marburg, Germany, Federal Republic

Braun, Konrad, Ebsdorfergrund, Germany, Federal

Republic of

Behringwerke Akitengesellschaft, Marburg, Germany, PATENT ASSIGNEE(S):

Federal Republic of (non-U.S. corporation)

NUMBER KIND DATE ______

US 5021243 19910604 PATENT INFORMATION:

US 1988-164486 19880304 (7) APPLICATION INFO.:

> NUMBER DATE _____

DE 1987-3707213 19870306 PRIORITY INFORMATION:

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

Robinson, Douglas W. PRIMARY EXAMINER:

ASSISTANT EXAMINER: Witz, Jean

Finnegan, Henderson, Farabow, Garrett, and Dunner LEGAL REPRESENTATIVE:

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

NUMBER OF DRAWINGS: 1 Drawing Figure(s); 1 Drawing Page(s) LINE COUNT: 254

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

A process for the preparation of a factor

VIII: C-deficient plasma is made available, in which a

starting plasma is consecutively treated with antibodies against von

Willebrand factor and antibodies against factor

VIII:Aq. The deficient plasma prepared in this way contains less than

0.5% residual activity of factor VIII:C.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 48 OF 72 USPATFULL on STN

9<u>1:1</u>247 USPATFULL ACCESSION NUMBER:

Lectin affinity chromatography of factor VIII TITLE: Tsay, Grace C., Walnut Creek, CA, United States INVENTOR(S):

Miles Inc., Elkhart, IN, United States (U.S. PATENT ASSIGNEE(S):

corporation)

NUMBER KIND DATE

US 4981951 19910101 US 1988-181001 19880414 (7) PATENT INFORMATION: <--

APPLICATION INFO.:

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: Moskowitz, Margaret ASSISTANT EXAMINER: Furman, Keith C.

LEGAL REPRESENTATIVE: Aston, David J., Bradley, Bertram

NUMBER OF CLAIMS: 6 EXEMPLARY CLAIM: 405 LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

A method of purifying a recombinant protein from a solution, such as AB tissue culture fluid, containing gylcoproteins. The affinity of lectins for specific glycoproteins is assessed and used to select a particular lectin specific for the contaminating glycoprotein(s). A sugar buffer such as alpha methyl mannoside prevents binding of the recombinant protein. The preferred lectin is lentil lectin, for use in separating recombinant Factor VIII from tissue culture fluid contaminated with rodent protein from the cell line used to produce the recombinant Factor VIII.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 49 OF 72 USPATFULL on STN L3

90:67728 USPATFULL ACCESSION NUMBER:

Method for purifying antihemophilic factor TITLE: Mathews, Rita W., New York, NY, United States INVENTOR(S):

Johnson, Alan J., New York, NY, United States

<--

New York University, New York, NY, United States (U.S. PATENT ASSIGNEE(S):

corporation)

NUMBER KIND DATE ______

US 4952675 PATENT INFORMATION: 19900828 19881229 (7) APPLICATION INFO.: US 1988-291516

20050510 DISCLAIMER DATE:

Continuation of Ser. No. US 1987-122372, filed on 19 RELATED APPLN. INFO.: Nov 1987, now patented, Pat. No. US 4847362 which is a continuation of Ser. No. US 1985-697267, filed on 1 Feb

1985, now patented, Pat. No. US 4743680

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: Nutter, Nathan M. LEGAL REPRESENTATIVE: Darby & Darby

11 NUMBER OF CLAIMS: EXEMPLARY CLAIM:

2 Drawing Figure(s); 2 Drawing Page(s) NUMBER OF DRAWINGS:

LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Disclosed are methods for purifying proteins including AHF by column chromatography in the presence of additives including sugars and polyhydric alcohols which serve to increase the electrostatic forces on the surface of said proteins while decreasing the hydrophobicity of said proteins resulting in preparations of such proteins of high purity and/or resolution and/or recovery.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L3 ANSWER 50 OF 72 WPIDS COPYRIGHT 2005 THE THOMSON CORP on STN

ACCESSION NUMBER: 1990-255000 [34] WPIDS

DOC. NO. NON-CPI: N1990-197573 DOC. NO. CPI: C1990-110404

TITLE: Production of von Willebrand factor

concentrates - by cleaving factor V111 C complex

on anion exchanger.

DERWENT CLASS: B04 D16 P34

INVENTOR(S): HEIMBURGER, N; KUMPE, G; WELLNER, K

PATENT ASSIGNEE(S): (BEHW) BEHRINGWERKE AG; (CENT-N) CENTEON PHARMA GMBH;

(AVET) AVENTIS BEHRING GMBH

COUNTRY COUNT: 2

PATENT INFORMATION:

PAT	TENT NO		KIN	ID DATE	WEEK	LA	PG
DE	3904354		 А	19900816	(199034)	·	 5<
ΕP	383234		Α	19900822	(199034)		<
	R: AT BE	СН	DE	ES FR GB	GR IT LI	LU NL	SE
ΑU	9049339		Α	19900830	(199042)		<
PT	93128		Α	19900831	(199043)		<
CA	2009946		Α	19900814	(199044)	•	<
JP	02264799		Α	19901029	(199049)		<
AU	638969		В	19930715	(199335)		<
ΕP	383234		В1	19941130	(199501)	GE	8<
	R: AT BE	CH	DE	DK ES FR	GB GR IT	LI LU	NL SE
DE	59007785		G [.]	19950112	(199507)		<
ES	2066020		Т3	19950301	(199515)		<
ΙE	65920		В	19951129	(199606)	•	<
JP	2930243		B2	19990803	(199936)		6<
KR	149999		В1	19980817	(200022)		<
CA	2009946		С	20000411	(200035)	EN	<
US	6239261		В1	20010529	(200132)		<

APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
DE 3904354	A	DE 1989-3904354	19890214
EP 383234	A	EP 1990-102716	19900212
JP 02264799	A	JP 1990-29777	19900213
AU 638969	В	AU 1990-49339	19900213
EP 383234	B1	EP 1990-102716	19900212
DE 59007785	G	DE 1990-507785	19900212
		EP 1990-102716	19900212
ES 2066020	Т3	EP 1990-102716	19900212
IE 65920	В	IE 1990-511	19900213
JP 2930243	B2	JP 1990-29777	19900213
KR 149999	B1	KR 1990-1674	19900212
CA 2009946	С	CA 1990-2009946	19900213
US 6239261	B1 Cont of	US 1990-478640	19900212
	Cont of	US 1991-759983	19910916
	Cont of	· US 1992-899936	19920617
		US 1994-253232	19940602

FILING DETAILS:

PATENT NO KIND

PATENT NO

```
AU 638969
                   B Previous Publ.
                                       AU 9049339
                   G Based on EP 383234
     DE 59007785
     ES 2066020 T3 Based on
                                       EP 383234
     JP 2930243
                   B2 Previous Publ. JP 02264799
PRIORITY APPLN. INFO: DE 1989-3904354
                                          19890214
     1990-255000 [34]
                       WPIDS
         3904354 A UPAB: 19930928
     Production of von Willestrand factor (vWF) concentrates from
     starting materials containing vWF as a complex with factor (
     VIII:C is effected by (a) preparing a solution containing the
     starting material and 5-30 weight % of a carbohydrate in an amino acid buffer
     with a pH of 5.5-6.5 and (b) treating the solution with an anion exchanger
     capable of binding factor (VIII):c thereby
     obtg. a VWF solution
          USE/ADVANTAGE - The concentrates are useful for treating
     von Willebrand's syndrome. The process gives high yields of high-purity
     vWF solns. which may be pasteurised to inactivate viruses.
     0/0
ABEO EP
          383234 B UPAB: 19950110
     A process for the preparation of pasteurised von
     Willebrand factor concentrate, which comprises
     a solution which contains von Willebrand factor (vWF)
     as complex with V III:C in a buffer of pH 5.5 to 6.5, which contains
     calcium and amino acids and has a carbohydrate concentration of 5-30% w/w,
     being treated with an anion exchanger to which F VIII:C binds, and the von
     Willebrand factor concentrate being obtd. from
     Dwg.0/0
     ANSWER 51 OF 72 BIOTECHDS COPYRIGHT 2005 THE THOMSON CORP. on STN
ACCESSION NUMBER: 1990-09347 BIOTECHDS
                  Recombinant Factor-VIII production - cloning, clinicals, and
TITLE:
                  commercial production;
                       Factor-VIII:c and human
                     recombinant von Willebrand factor
                     preparation by mammal cell culture in serum-free
                     culture medium; process monitoring, systems control and
                     validation (conference abstract)
                  Adamson S R
CORPORATE SOURCE: Genetics-Inst.
                  Genetics Institute, One Burtt Road, Andover, MA 01810, USA.
LOCATION:
                 Abstr.Pap.Am.Chem.Soc.; (1990) 199 Meet., Pt.1,
SOURCE:
                 BIOT80
                 CODEN: ACSRAL
DOCUMENT TYPE:
                 Journal
LANGUAGE:
                 English
      1990-09347 BIOTECHDS
      Hemophilia A is a sex-linked bleeding disorder caused by a deficiency or
      abnormality of the blood-clotting protein Factor-VIII
      :c. The available therapy (plasma concentrate) is.
      costly and associated with a finite risk of viral infections. The
      molecular cloning of Factor-VIII:c signaled
      the provision of pure Factor-VIII:c.
      Subsequently, Factor-VIII:c was produced by
      recombinant mammal cells in the presence of high concentrations (10-20%
      v/v) of animal serum, but not in serum-free culture media. The essential
      component of serum was identified as von Willebrand
      factor (vWF). The requirement for serum can be eradicated by the
      addition of high concentrations of phospholipid vesicles to the culture
      medium, or by the co-expression of human vWF and Factor-
      VIII:c from the same recombinant cell. Development of
      these techniques was described with respect to strategies used in process
      monitoring, control and validation. These strategies are essential in
      order to address scientific and regulatory requirements or concerns
      directed towards assuring the production of high quality, safe products.
```

ΑB

AN

AB

(1 ref)

ANSWER 52 OF 72 CAPLUS COPYRIGHT 2005 ACS on STN DUPLICATE 2

ACCESSION NUMBER: 1989:530280 CAPLUS

DOCUMENT NUMBER: 111:130280

Fluoroplastic immunoaffinity columns for purification

of blood proteins

INVENTOR(S): Zimmerman, Theodore S.; Fulcher, Carol A. Scripps Clinic and Research Foundation, USA PATENT ASSIGNEE(S):

SOURCE: U.S., 5 pp. CODEN: USXXAM

DOCUMENT TYPE: Patent English LANGUAGE:

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

TITLE:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 4831118	Α	19890516	US 1987-83670	19870807 <
PRIORITY APPLN. INFO.:			US 1987-83670	19870807

A protein purification column comprises a fluoroplastic substrate matrix having low reactivity to proteins, said matrix being capable of maintaining monoclonal antibodies attached thereto in an external configuration and preventing interaction with the protein to be bound to the antibody, and a monoclonal antibody attached to the substrate, the monoclonal antibody having a specific affinity for the protein to be isolated. Specific protein is isolated and purified from a solution by: (1) attaching protein-specific monoclonal antibody to the fluoroplastic substrate matrix to form an antibody-substrate conjugate; and (2) contacting protein to be isolated, in an appropriate buffer solution, with the antibody-substrate conjugate. An appropriate buffer may be applied to remove nonantibody-bound contaminants, followed by an appropriate eluting agent to remove the protein from the monoclonal antibody. Blood-coagulation factor VIII was purified from reconstituted Armour Factorate conc . by affinity chromatog. on perfluorocarbon support containing .apprx.2 mg anti-von Willebrand Factor monoclonal antibody 2.2.9/mL. The factor was eluted from the column with 0.35 M CaCl2 buffer. Total protein recovery was 95% (85% in eluate, 10% in pass-through and wash) when a slow flow rate (2.2 mL/min loading, 2.3 and 0.7 mL/min washing) was used.

ANSWER 53 OF 72 USPATFULL on STN.

89:56511 USPATFULL ACCESSION NUMBER:

Method for purifying antihemophilic factor TITLE:

Mathews, Rita W., both New York, NY, United States Johnson, Alan J., both New York, NY, United States INVENTOR(S):

New York University, New York, NY, United States (U.S. PATENT ASSIGNEE(S):

corporation)

NUMBER	KIND	DATE

US 4847362 19890711 <--PATENT INFORMATION:

US 1987-122372 19871119 (7) APPLICATION INFO.:

Continuation of Ser. No. US 1985-697267, filed on 1 Feb RELATED APPLN. INFO.:

1985, now patented, Pat. No. US 4743680

DOCUMENT TYPE: Utility Granted FILE SEGMENT: Kight, John PRIMARY EXAMINER: ASSISTANT EXAMINER: Nutter, Nathan M. LEGAL REPRESENTATIVE: Darby & Darby

NUMBER OF CLAIMS: 23 EXEMPLARY CLAIM:

NUMBER OF DRAWINGS: 3 Drawing Figure(s); 3 Drawing Page(s)

1215 LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The present invention relates to a high-recovery, high-resolution method AB for purifying antihemophilic factor by using column chromatography techniques in the presence of sugars, polyhydric alcohols, amino acids or salts.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 54 OF 72 USPATFULL on STN L3ACCESSION NUMBER: 89:1286 USPATFULL

Phospholipid affinity purification of Factor TITLE:

INVENTOR(S): Brown, James E., Lafayette, CA, United States

Cowgill, Cynthia A., Berkeley, CA, United States

Miles Laboratories, Inc., Elkhart, IN, United States PATENT ASSIGNEE(S):

(U.S. corporation)

NUMBER KIND DATE -----

US 4795806 19890103 PATENT INFORMATION: <--

19870716 (7) US 1987-74123 APPLICATION INFO.:

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

Schain, Howard E. PRIMARY EXAMINER: ASSISTANT EXAMINER: Kushan, Jeff P.

LEGAL REPRESENTATIVE: Aston, David J., Simonton, Pamela A.

NUMBER OF CLAIMS: 11 EXEMPLARY CLAIM: 1 449 LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Affinity purification of Factor VIII:C,

both plasma derived and genetically engineered, using coupled

phosphatidylserine (PS) as the predominant phospholipid (PH) results in

a high degree of purity of Factor VIII:C,

similar to that previously demonstrated with monoclonal antibodies

specific to either Factor VIII:C or von

Willebrand factor. Phospholipids that can be used in combination with PS are phosphatidycholine (PC) and

phosphatidylethanolamine (PE).

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 55 OF 72 WPIDS COPYRIGHT 2005 THE THOMSON CORP on STN

ACCESSION NUMBER: 1990-007450 [01] WPIDS

C1990-003204 DOC. NO. CPI:

Separation of plasma proteins, especially factor-VIII - by TITLE:

chromatography on moderately ionic anion-exchange resin.

DERWENT CLASS: B04 J01

BURNOUF, M; BURNOUF, T; BURNOUF-RADOSEVICH, M INVENTOR(S):

PATENT ASSIGNEE(S): (REGI-N) CENT REGIONAL TRANSFUSION SANGUINE; (TRAN-N)

CENT REGION TRANSFU; (REGI-N) CENT REGIONAL TRANS;

(TRAN-N) CENT REG TRANSFUSION SANGUINE LILLE

COUNTRY COUNT: 22

PATENT INFORMATION: DAMENIA NO

PAT	TENT NO	KIND DATE	WEEK	, LA	PG
WO	8912065	A 19891214	(199001)	FR	19<
	RW: AT BE CH	DE FR GB IT	LU NL SE		
	W: AU DK FI	JP KR NO SU	US		
FR	2632309	A 19891208	(199005)		<
ΑU	8930682	A 19900105	(199012)		<
EP	359593	A 19900321	(199012)	FR	<
	R: AT BE CH	DE ES FR GB	GR IT LI	LU NL	SE
FI	9000397	A 19900125	(199018)		<
NO	9000529	A 19900507	(199024)		<
DK	9000299	A 19900328	(199027)		<
JΡ	03501974	W 19910509	(199125)		<
ΑU	9211383	A 19920514	(199228)		<
US	5252709	A 19931012	(199342)		5<
SU	1837880	A3 19930830	(199519)		6<
EP	359593	B1 19950426	(199521)	FR	10<
	R: AT BE CH	DE ES FR GB	GR LT LI	LU NL	SE

NO	177188	В	19950424	(199522)		<
DE	68922358	E	19950601	(199527)		<
ES	2070919	Т3	19950616	(199531)		<
ΓI	96210	В	19960215	(199611)		<
JP	2805364	B2	19980930	(199844)		6<
KR	9710923	В1	19970702	(199946)		<
CA	1340742	С	19990914	(200004)	FR	<
ΕP	359593	B2	20040107	(200405)	FR	
	R: AT BE CH	DE	ES FR GB	GR IT LI	LU NL	SE
DK	2004000872	Α	20040603	(200443)		
DK	175322	В	20040823	(200456)		

APPLICATION DETAILS:

PAT	ENT NO	KIND	APPLICATION .	DATE
WO	8912065	A	WO 1989-FR50	19890208
FR	2632309	A	FR 1988-7530	19880607
ΕP	359593	A	EP 1989-400348	19890208
JP	03501974	W	JP 1989-502342	19890208
ΑU	9211383	A	AU 1992-11383	19920303
		Div ex	AU 1989-30682	
US	5252709	A	WO 1989-FR50	19890208
			US 1990-460972	19900406
SU	1837880	A3	WO 1989-FR50	19890208
			SU 1990-4743107	19900206
ΕP	359593	B1	EP 1989-400348	19890208
ИО	177188	В	WO 1989-FR50	19890208
			NO 1990-529	19900205
DE	68922358	E	DE 1989-622358	19890208
			EP 1989-400348	19890208
ES	2070919	Т3	EP 1989-400348	19890208
FI	96210.	В	WO 1989-FR50	19890208
			FI 1990-397	19900125
JΡ	2805364	B2	JP 1989-502342	19890208
			WO 1989-FR50	19890208
KR	9710923	B1	WO 1989-FR50	19890208
			KR 1990-700239	19900206
	1340742	С	CA 1989-590961	19890214
	359593	B2	EP 1989-400348	19890208
	2004000872	A	DK 2004-872	20040603
DK	175322	В	WO 1989-FR50	19890208
			DK 1990-299	19900206

FILING DETAILS:

PATENT NO	KIND	PATENT NO	
US 5252709	A Based on	WO 8912065	
NO 177188	B Previous Publ.	NO 9000529	
DE 68922358	E Based on	EP 359593	
ES 2070919	T3 Based on	EP 359593	
JP 2805364	B2 Previous Publ.	JP 03501974	
	Based on	WO 8912065	
DK 175322	B Previous Publ.	DK 9000299	

PRIORITY APPLN. INFO: FR 1988-7530 19880607

1990-007450 [01] WPIDS AB

WO 8912065 A UPAB: 19960520

Separation of human or animal plasma proteins is effected by chromatographing a solubilised cryoprecipitate fraction on a moderately ionic anion-exchange resin, such that very large mols. may be retained and hydrophobic interactions come into play, and recovering each of the proteins selectively by increasing the ionic strength of the elution buffer.

USE/ADVANTAGE - The process is capable of producing a factor VIII concentrate with a specific activity above 100 IU/mg, useful for treating type-A haemophilia, as well as fibrinogen, von Willebrand factor (vWF) and fibronectin concentrates requiring

further purificn.

Dwg.0/0 Dwg.0/0

ABEQ US 5252709 A UPAB: 19931202

Factor VIII is sepd. from plasma also contg. von willebrand's factor (VWF), fibrinogen and fibronectin by (A) chromatographing a buffer and a solubilised fraction of cryoprecipitated plasma on an exchange column having a DEAE gp. fixed to a vinyl polymer type gel, i.e. Fractogel TSK, which absorbs factor VIII,VWF and fibronectin and allows fibrinogen to pass into a 1st eluate, (B) increasing the ionic strength of the buffer to allow fibronectin and VWF to pass into a 2nd eluate and (C) further increasing the ionic strength of the buffer to allow elution of factor VIII.

The specific activity of **factor VIII:C** in the initial fraction is at least 0.1 UV.mg. The initial fraction has pref. been pre-purified by treatment with Al(OH)3, cooling to 14-16 deg. C, centrifuging and recovery of the superntant. The buffer contains 2-4 g/ltr lysine and 8-11 g/ltr glycine and 0.11M NaCl. The ionic strength in (B) is increased by raising the NaCl concentration to 0.15 M and in (C) to 0.25. A virus inactivating treatment is also carried out on the plasma prior to chromatographing.

ADVANTAGE - The desired protein can be separated using a single chromatography column avoiding expensive and cumbersome further purification, which can decrease the activity of the protein, VWF, fibronectin and fibrinogen can be recovered separately.

Dwg. 0/0

ABEQ EP 359593 B UPAB: 19950602

Process for the separation of human or animal plasma proteins and for the preparation of concentrates of the said proteins for therapeutic use, characterised in that it comprises the following steps: the cryoprecipitate fraction of plasma, consisting essentially of fibrinogen, fibronectin, Willebrand's factor and Factor VIII, is used as starting material; the cryoprecipitate, dissolved in aqueous solution, is subjected to a single separation by chromatography on an anion exchange resin of which the matrix is a gel of the macroreticular vinyl polymer type which, as a result of its hydrophobic and porous properties, is capable of retaining the Factor VIII-Willebrand's factor complex; and the different proteins are recovered selectively by successive increases in the ionic strength of the elution buffer.

Dwg.0/0

L3 ANSWER 56 OF 72 CAPLUS COPYRIGHT 2005 ACS on STN DUPLICATE 3

ACCESSION NUMBER: 1989:612495 CAPLUS

DOCUMENT NUMBER: 111:212495

TITLE: Comparison of the in vitro characteristics of von

Willebrand factor in British and commercial factor VIII concentrates

AUTHOR(S): Lawrie, A. S.; Harrison, P.; Armstrong, A. L.;

Wilbourn, B. R.; Dalton, R. G.; Savidge, G. F.

CORPORATE SOURCE: Rayne Inst., St. Thomas' Hosp., London, SE1 7EH, UK

British Journal of Haematology (1989),

73(1), 100-4

CODEN: BJHEAL; ISSN: 0007-1048

DOCUMENT TYPE: Journal LANGUAGE: English

SOURCE:

AB Qual./quant. anal. of von Willebrand factor antigen
(vWf:Ag) in either heat or solvent/detergent-treated factor VIII
concs., used for hemophilia replacement therapy, was undertaken to
assess their suitability for the treatment of von Willebrand's disease
(vWD). For the 1st time, immunoaffinity purified vWf:Ag (Monoclate
byproduct) was also evaluated by in vitro assessment. Potencies of vWf:Ag
varied considerably but were consistently higher (28.9-420.5 iu/mL) than
factor VIII:C (one-stage) activity (8.13-42.44
iu/mL). The functional activity of vWf was assessed by either ristocetin
cofactor (vWf:RCo) or collagen binding methods (vWf:CBA) with typical
vWf:RCo/vWf:Ag ratios ranging from 0.08 to 0.94. Multimeric anal.
confirmed that in vitro biol. activity was dependent on the presence of

the high mol. weight forms of vWf:Ag. A significant correlation between

vWf:RCo activity and collagen binding was observed in all of the concs. with the exception of the immunopurified product.

Apparently, either NHS 8Y (mean wWfRCo/vWf:Ag = 0.9), Haemate P (mean vWf:RCo/vWf:Ag = 0.69), and high purity Octapharma V.I (vWf:RCo/vWf:Ag = 0.82) which contain medium/high mol. weight vWf:Ag multimers are likely to be the most cost-effective in the treatment of symptomatic severe vWD patients than other currently available concs.

L3 ANSWER 57 OF 72 USPATFULL on STN

ACCESSION NUMBER: 88:29502 USPATFULL

TITLE: Method for purifying antihemophilic factor
INVENTOR(S): Mathews, Rita W., New York, NY, United States
Johnson, Alan J., New York, NY, United States

PATENT ASSIGNEE(S): New York University, New York, NY, United States (U.S.

corporation)

NUMBER KIND DATE

PATENT INFORMATION: US 4743680 19880510 <--

APPLICATION INFO.: US 1985-697267 19850201 (6)
DOCUMENT TYPE: Utility

FILE SEGMENT: Granted
PRIMARY EXAMINER: Kight, John
ASSISTANT EXAMINER: Nutter, Nathan M.
LEGAL REPRESENTATIVE: Darby & Darby

NUMBER OF CLAIMS: 24 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 3 Drawing Figure(s); 3 Drawing Page(s)

LINE COUNT: 1233

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Disclosed are methods for purifying proteins including AHF by column chromatography in the presence of additives including sugars and polyhydric alcohols which serve to increase the electrostatic forces on the surface of said proteins while decreasing the hydrophobicity of said proteins resulting in **preparations** of such proteins of high purity and/or resolution and/or recovery.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L3 ANSWER 58 OF 72 MEDLINE ON STN ACCESSION NUMBER: 89187703 MEDLINE DOCUMENT NUMBER: PubMed ID: 3149046

TITLE: Factor VIII procoagulant protein interacts with

phospholipid vesicles via its 80 kDa light chain.

AUTHOR: Kemball-Cook G; Edwards S J; Sewerin K; Andersson L O;

Barrowcliffe T W

CORPORATE SOURCE: , National Institute for Biological Standards and Control,

Potters Bar, Herts, UK.

SOURCE: Thrombosis and haemostasis, (1988 Dec 22) 60 (3)

442-6.

Journal code: 7608063. ISSN: 0340-6245.

PUB. COUNTRY: GERMANY, WEST: Germany, Federal Republic of

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 198905

ENTRY DATE: Entered STN: 19900306

Last Updated on STN: 19900306 Entered Medline: 19890501

In a previous report, we detailed fractionation of polyclonal human antiFactor VIII:C into a component directed
exclusively against the phospholipid-binding site on Factor VIII (PL-site
antibody) and another directed at other sites (non-PL-site antibody). The
location on the F.VIII molecule of its PL-binding site has now been
studied by two different methods using this fractionated 125I-labelled
anti-F.VIII:C Fab'. The first method was modified from that of Weinstein
et al. (Proc Natl Acad Sci USA 1981; 78: 5137-41), involving
electrophoresis of F.VIII peptide-125I-Fab' A/F.VIII immunocomplexes in

SDS-polyacrylamide gels. PL-site antibody reacted with F.VIII peptides of apparent Mr approximately 80 kDa and sometimes 160 kDa in plasma and concentrate, but not with larger peptides. Non-PL-site antibody, however, reacted with a range of peptides of apparent Mr 90 kDa to 280 kDa. In addition, when purified F.VIII containing heavy and light chains (HC + LC), and isolated LC peptides were analysed, PL-site antibody bound to LC peptides whereas non-PL-site antibody did not. The second method used the antibody pools in immunoradiometric assays (IRMA's) of purified F.VIII peptides. Both labels measured similar amounts of F.VIII:Ag in a sample of purified F.VIII containing both HC and LC; on assaying an HC preparation, however, PL-site label measured only 2% of F.VIII:Ag found by non-PL-site label, indicating that PL-binding sites are absent in HC preparations. These results indicate that F.VIII binds to PL via its 80 kDa light chain.

L3 ANSWER 59 OF 72 USPATFULL on STN

ACCESSION NUMBER: 87:45187 USPATFULL

TITLE: Tsolation of human plasma procoagulant protein factor

VIII from biological factors

INVENTOR(S): Herring, Steven W., San Dimas, CA, United States

PATENT ASSIGNEE(S): Alpha Therapeutic Corporation, Los Angeles, CA, United

States (U.S. corporation)

NUMBER KIND DATE

PATENT INFORMATION: US 4675385 19870623 <--

APPLICATION INFO.: US 1985-716456 19850327 (6)

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: Schain, Howard E.

LEGAL REPRESENTATIVE: Christie, Parker & Hale

NUMBER OF CLAIMS: 25 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 3 Drawing Figure(s); 3 Drawing Page(s)

LINE COUNT: 979

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB A rapid and simple process for purifying human, bovine and porcine

procoagulant protein Factor VIII on a large scale using sequential high performance size exclusion chromatography under, first, low salt concentration conditions and, second, under high salt concentration conditions from reconstituted commercial Factor VIII :C (complexed Factor VIII) concentrate. The chromatographic separation is carried out on a high performance size exclusion chromatographic column packed with porous beads having a particle size of from about 13 to about 35 microns, pore diameters of from about 500 to about 2000 Angstroms and a pore volume of from about 1.0 to about 1.8 ml per gram. The first chromatographic separation is carried in a buffered aqueous solution using the buffered aqueous solution as an eluant. The low molecular weight constituents (impurities) are separated from Factor VIII and the high molecular weight constituents (impurities). A second chromatographic separation may be carried out after Factor VIII has been dissociated in a buffered solution having a concentration of from about 0.25 to about 0.45M calcium ion. The second chromatographic column is packed with some packing as the first column and is eluted with a buffered aqueous

can be purified in less than two hours. The process is amenable to scale up.

solution containing 0.25 to 0.45M calcium ion. In a column of

2.5+60 cm, 4 gms of commercial Factor VIII concentrate

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L3 ANSWER 60 OF 72 USPATFULL on STN

ACCESSION NUMBER: -87:39834 USPATFULL

TITLE: Isolation and culture of adrenal medullary endothelial

cells producing blood clotting factor

VIII:C

INVENTOR(S): Pollard, Harvey B., Potomac, MD, United States

Ornberg, Richard, Bethesda, MD, United States Banerjee, Dipak, Rockville, MD, United States Youdim, Moussa, Rockville, MD, United States Lelkes, Peter, Rockville, MD, United States Heldman, Eli, Rockville, MD, United States

PATENT ASSIGNEE(S):

The United States of America as represented by the Secretary of the Department of Health and Human Services, Washington, DC, United States (U.S.

government)

KIND NUMBER DATE _______

US 4670394 PATENT INFORMATION: 19870602 <--19841116 (6) APPLICATION INFO .: US 1984-672451

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

Wiseman, Thomas G. PRIMARY EXAMINER: ASSISTANT EXAMINER: Maurey, Karen LEGAL REPRESENTATIVE: Holman & Stern

NUMBER OF CLAIMS: 10 EXEMPLARY CLAIM:

NUMBER OF DRAWINGS: 6 Drawing Figure(s); 5. Drawing Page(s)

LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The present invention discloses a new line of endothelial cell of adrenal medullary origin capable of producing blood clotting

Factor VIII: C. A method of isolating and

culturing said cell line has also been disclosed. Factor

VIII:C is useful in treating hemophilia.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 61 OF 72 EMBASE COPYRIGHT 2005 ELSEVIER INC. ALL RIGHTS RESERVED.

on STN

TITLE:

87115012 EMBASE ACCESSION NUMBER:

1987115012 DOCUMENT NUMBER:

Characteristics of a heat treated antihaemophilic

cryoprecipitate.

Skjonsberg O.H.; Gravem K.; Kierulf P.; Godal H.C. AUTHOR:

CORPORATE SOURCE: Haematological Research Laboratory and Central Laboratory,

Ulleval Hospital, 0407 Oslo 4, Norway

Thrombosis Research, (1987) Vol. 45, No. 5, pp. 625-634. SOURCE:

CODEN: THBRAA

COUNTRY: United States

DOCUMENT TYPE: Journal

037 Drug Literature Index FILE SEGMENT:

025 Hematology 022 Human Genetics Pharmacology 030

LANGUAGE: English

ENTRY DATE: Entered STN: 911211

Last Updated on STN: 911211

In order to evaluate the influence of heat treatment (68°C for 24 AB or 72 hours) on the essential components of antihaemophilic

cryoprecipitate, i.e. factor VIII coagulant activity (VIII:C), von

Willebrand factor (VIIIR: Ag and VIIIR: RCF) and

fibrinogen, ordinary lyophilized cryoprecipitate was compared to heat treated, aminoacid-enriched specimens. The median reduction in

factors VIII:C, VIIIR:Ag, VIIIR:RCF and fibrinogen during lyophilization of ordinary cryoprecipitate was 26 per cent, 11 per cent, 1 per cent and 8.5 per cent, respectively. Heat treatment of such cryoprecipitate resulted in 85 to 98.5 per cent reduction in these parameters, while the reduction following lyophilization and heat treatment (24 hours) of aminoacid-containing preparations was not significantly different from non-heated, ordinary cryoprecipitate. Following heating of aminoacid-enriched cryoprecipitate for 72 hours, only factor VIIIR: RCF was significantly reduced (32.5 per cent) compared to non-heated samples. Ordinary

cryoprecipitate was almost insoluble following heat treatment. Enrichment with aminoacids, however, made the heat treated cryoprecipitate fully soluble, but the content of these vials were slightly slower in dissolving than non-heated preparations. Ultracentrifugation prior to lyophilization and heating did not improve the solubility. treatment proves to be efficient in inactivating viral agents, we conclude that heated (68°C for 24 hours), aminoacid-enriched cryoprecipitate may be a convenient product for treating haemophilia A, von Willebrand's disease and hypofibrinogenemia.

ANSWER 62 OF 72 USPATFULL on STN

86:55170 USPATFULL ACCESSION NUMBER:

Deglycosylated Human Factor VIII: TITLE:

Chavin, Stephen I., Rochester, NY, United States INVENTOR(S):

Fay, Philip J., Rochester, NY, United States

PATENT ASSIGNEE(S): University of Rochester, Rochester, NY, United States

(U.S. corporation)

NUMBER KIND DATE

PATENT INFORMATION: US 1984-570728 Continuation 19860930

19840113 (6) APPLICATION INFO.:

RELATED APPLN. INFO.: Continuation-in-part of Ser. No. US 1982-405456, filed

on 5 Aug 1982, now patented, Pat. No. US 4495175

DOCUMENT TYPE: Utility Granted FILE SEGMENT: PRIMARY EXAMINER: Kight, John ASSISTANT EXAMINER: Nutter, Nathan M.

LEGAL REPRESENTATIVE: Hallenbeck, Robert M., LuKacher, Martin L., Gibblin,

James

NUMBER OF CLAIMS: 2 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 8 Drawing Figure(s); 6 Drawing Page(s)

635 LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Highly purified, biologically active Human Factor VIII :C having specific activities of about 4000-8000 units per milligram of protein is prepared. In the method of preparation , an AHF concentrate is solubilized or equilibrated in an aqueous medium and treated to change the effective Stokes' radius of the Factor VIII:C to an apparently low value and

then subjected to a separation from the concentrate. Treatment

of the highly purified Factor VIII:C with

a mixture of glycosidases causes substantial removal of carbohydrate side chains without reduction of procoagulant activity and with retention of significant in vivo survival time.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 63 OF 72 USPATFULL on STN

ACCESSION NUMBER: 86:16911 USPATFULL

TITLE: _Purification of factor VIII with insoluble matrix

having free sulfate groups covalently bound thereto

Saundry, Richard H., London, England INVENTOR(S):

Savidge, Geoffrey F., Kent, England

The Special Trustees for St. Thomas' Hospital, London, PATENT ASSIGNEE(S):

England (non-U.S. corporation)

NUMBER KIND DATE -----

US 4578218 US 1985-699957 PATENT INFORMATION: 19860325 <--

19850208 (6) APPLICATION INFO.:

> NUMBER DATE ______

GB 1984-3473 19840209 PRIORITY INFORMATION:

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

PRIMARY EXAMINER: Schain, Howard E.

LEGAL REPRESENTATIVE: Cushman, Darby & Cushman

NUMBER OF CLAIMS: 11 EXEMPLARY CLAIM: 1 LINE COUNT: 765

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Factor VIII is purified by adsorption onto a insoluble matrix having free sulphate groups, such as dextran sulphate, and selective elution therefrom.

A suitable eluant for the purification of the von Willebrand protein (Factor VIIIR:vWp) is citrate buffer, pH 6.85, containing 0.47 M sodium chloride and 2.14 mM calcium chloride.

A suitable eluant for the purification of the Factor VIII complex (Factor VIIIR: Ag, Factor VIIIR: vWp and Factor VIII: c) is citrate buffer at pH value between 6.2 and 7.3 containing

1.0 M glycine, 2.14 mM calcium chloride and 0.5 M sodium chloride at +4° C.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L3 ANSWER 64 OF 72 IFIPAT COPYRIGHT 2005 IFI on STN DUPLICATE 4

01572348 IFIPAT; IFIUDB; IFICDB AN

PREPARATION OF HIGHLY PURIFIED HUMAN TITLE:

ANTIHEMOPHILIC FACTOR

Chavin, Stephen I, Rochester, NY INVENTOR(S):

Fay, Philip J, Rochester, NY

PATENT ASSIGNEE(S):

University of Rochester, Rochester, NY

PRIMARY EXAMINER:

AGENT:

Rosen, Sam Aston, David J

Leitereg, Theodore J

PK DATE NUMBER

PATENT INFORMATION: US 4495175 A 19850122

(CITED IN 021 LATER PATENTS)

APPLICATION INFORMATION: US 1982-405456 19820805

5 Aug 2002 EXPIRATION DATE:

US 4495175 19850122 FAMILY INFORMATION:

DOCUMENT TYPE: Utility FILE SEGMENT: CHEMICAL GRANTED

MICROFILM REEL NO: 004119 FRAME NO: 0680

NUMBER OF CLAIMS: 10

GRAPHICS INFORMATION: 3 Drawing Sheet(s), 5 Figure(s).

Highly purified, biologically active Human Antihemophilic Factor (AHF)

preparations are prepared having specific activities of about

4000-8000 units per milligram of AHF. In the method of

preparation an AHF concentrate, prepared by fractionation of plasma to partially remove fibrinogen, fibronectin and other plasma components is subjected to a separation on the basis of Stokes' radius to separate AHF from the bulk of remaining proteins in the AHF concentrate. The pooled fractions containing AHF activity are concentrated by precipitation with ammonium sulfate, sodium sulfate, etc., by diafiltration, by PEG addition, or the like. The concentrate, is solubilized or equilibrated in an aqueous medium and treated to change the effective Stokes' radius of the AHF to an apparently low value and then subjected to a separation from the concentrate. The AHF pool from above is treated to remove cations by dialysis against an appropriate buffer of lower ionic strength and chromatographed on an anionexchange medium. The AHF fraction from the above chromatography, is a highly purified AHF preparation.

CLMN

GΙ 3 Drawing Sheet(s), 5 Figure(s). ANSWER 65 OF 72 USPATFULL on STN

85:61585 USPATFULL ACCESSION NUMBER:

TITLE: Ultrapurification of factor VIII using monoclonal

antibodies

INVENTOR(S): Zimmerman, Theodore S., La Jolla, CA, United States

Fulcher, Carol A., La Jolla, CA, United States

PATENT ASSIGNEE(S): Scripps Clinic and Research Foundation, La Jolla, CA,

United States (U.S. corporation)

NUMBER KIND -----

US 32011 PATENT INFORMATION: 19851022

19821130 (Original) 19831221 (6) 19811214 (Original) US 4361509

US 1983-563795 APPLICATION INFO.:

US 1981-330105

DOCUMENT TYPE: Reissue Granted FILE SEGMENT:

Schain, Howard E. PRIMARY EXAMINER:

NUMBER OF CLAIMS: 36 EXEMPLARY CLAIM: 30 LINE COUNT: 715

CAS INDEXING IS AVAILABLE FOR THIS PATENT. CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 66 OF 72 MEDLINE on STN ACCESSION NUMBER: 85207837 MEDLINE DOCUMENT NUMBER: PubMed ID: 3922990

Standardization of factor VIII: establishment and use of TITLE:

secondary standards.

Panicucci F; Angeloni G; Arrighi S; Bucci E; DiMambro G; AUTHOR:

Lecchini L; Pitruzzello S; Positano M

SOURCE: Journal of biological standardization, (1985 Apr)

13 (2) 115-21.

Journal code: 0400335. ISSN: 0092-1157.

PUB. COUNTRY: ENGLAND: United Kingdom
DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

English LANGUAGE:

FILE SEGMENT: Priority Journals

198507 ENTRY MONTH:

ENTRY DATE: Entered STN: 19900320

> Last Updated on STN: 19900320 Entered Medline: 19850725

Two secondary standards for use in routine assays of Factor VIII in AΒ therapeutic concentrates and in patients, plasmas, respectively, have been established in a multicenter collaborative study. In order to assess the effect of the adoption of these preparations as common Secondary Standards a comparative assay has been performed: one sample of a Factor VIII concentrate of intermediate purity and one plasma sample have been tested in two laboratories for Factor VIII: c activity using as reference, among others, the common working standard. Analysis of the results shows that with the plasma sample the differences of the estimates obtained with any of the references in our two laboratories were not statistically significant (P greater than 0.3), while with the concentrate sample the differences were always statistically significant (P less than 0.005). The study shows that the adoption of common working standards (besides the uniformity in assay method, reagents and basic equipment) is not sufficient to eliminate interlaboratory variation in the measurement of Factor VIII:C.

ANSWER 67 OF 72 USPATFULL on STN

ACCESSION NUMBER: 84:51336 USPATFULL

Heparin polyelectrolyte polymer complex TITLE:

Johnson, John H., Kirkwood, MO, United States INVENTOR(S):

Monsanto Company, St. Louis, MO, United States (U.S. PATENT ASSIGNEE(S):

corporation)

PATENT INFORMATION: US 4471112 US 4471112 19840911 US 1983-460227 19830124 (6) 19840911

APPLICATION INFO.:

Division of Ser. No. US 1982-392929, filed on 28 Jun RELATED APPLN. INFO.:

1982, now patented, Pat. No. US 4397841

<--

<--

<--

DOCUMENT TYPE: Utility

FILE SEGMENT: Granted PRIMARY EXAMINER:

Brown, Johnnie R.

LEGAL REPRESENTATIVE: Meyer, Scott J., Williams, Jr., James W.

NUMBER OF CLAIMS: 4 EXEMPLARY CLAIM: LINE COUNT: 472

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

A concentrate of blood coagulation Factor

VIII:C is obtained in high yield by fractionation of

blood plasma with a sequence of adsorption steps employing two different water-insoluble, cross-linked polyelectrolyte copolymers, each in the

presence of exogenous heparin.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 68 OF 72 USPATFULL on STN

ACCESSION NUMBER: 83:34248 USPATFULL

Production of blood coagulation factor TITLE:

INVENTOR(S): Johnson, John H., Kirkwood, MO, United States

Monsanto Company, St. Louis, MO, United States (U.S. PATENT ASSIGNEE(S):

corporation)

NUMBER KIND DATE -----

PATENT INFORMATION:

US 4397841 19830809 US 1982-392929 19820628 (6)

DOCUMENT TYPE: Utility FILE SEGMENT: Granted PRIMARY EXAMINER: Rosen, Sam

LEGAL REPRESENTATIVE: Meyer, Scott J., Williams, Jr., James W.

NUMBER OF CLAIMS: EXEMPLARY CLAIM: 504 LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

A concentrate of blood coagulation Factor

VIII: c is obtained in high yield by fractionation of blood plasma with a sequence of adsorption steps employing two different water-insoluble, cross-linked polyelectrolyte copolymers, each in the presence of exogenous heparin.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 69 OF 72 USPATFULL on STN

ACCESSION NUMBER: 2:57885 USPATFULL

TITLE: Ultrapurification of factor VIII using monoclonal

antibodies

Zimmerman, Theodore S., La Jolla, CA, United States INVENTOR(S):

Fulcher, Carol A., La Jolla, CA, United States

PATENT ASSIGNEE(S): Scripps Clinic and Research Foundation, La Jolla, CA, United States (U.S. corporation)

NUMBER KIND DATE US 4361509 PATENT INFORMATION: 19821130

19811214 (6) APPLICATION INFO.: US 1981-330105

DOCUMENT TYPE: Utility Granted FILE SEGMENT:

PRIMARY EXAMINER: Schain, Howard E.

NUMBER OF CLAIMS: 16 EXEMPLARY CLAIM: 1 596 LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB A method of preparing high purity procoagulant protein comprising the steps of (a) adsorbing a VIII:C/VIII:RP complex from a plasma or commercial concentrate source of factor VIII onto agarose beads bound to a monoclonal antibody specific to VIII:RP, (b) eluting VIII:C with a salt solution, (c) adsorbing the eluted VIII:C on an animohexyl agarose column and eluting the VIII:C with a salt solution.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 70 OF 72 CAPLUS COPYRIGHT 2005 ACS on STN

ACCESSION NUMBER: 1982:213755 CAPLUS

DOCUMENT NUMBER: 96:213755

TITLE: Porcine factor VIII:C

prepared by affinity interaction with von

Willebrand factor and heterologous

antibodies: sodium dodecyl sulfate polyacrylamide gel

analysis

AUTHOR(S): Knutson, Gaylord J.; Fass, David N.

CORPORATE SOURCE: Mayo Clin. Mayo Found., Rochester, MN, 55905, USA

SOURCE: Blood (1982), 59(3), 615-24

CODEN: BLOOAW; ISSN: 0006-4971

DOCUMENT TYPE: Journal LANGUAGE: English

AB The concentration and partial purification of porcine factor VIII:

 ${f c}$ from plasma by conventional precipitation and chromatog. techniques was

reported. Blood from heparinized animal(s) was collected in citrate,

soybean trypsin inhibitor, e-aminocaproic acid, and benzamidine.

After Al(OH)3 adsorption, polyethylene glycol-6000 precipitation, QAE-cellulose

chromatog., and dextran sulfate-agarose chromatog, the factor

VIII: C conc. was 8100-fold purified with an

overall yield of 24%, and thrombin treatment of the factor

VIII: c gave an activation coefficient of up to 56. The

activation coefficient of plasma factor VIII:C

conc. on von Willebrand factor-agarose

produced a 2-fold increase in specific activity. This product was applied

to a 2nd affinity resin, the acidic IgG fraction of human antihuman

factor VIII:C coupled to agarose. The

inactive material which eluted at pH 2.8 from this column and from a similarly prepared nonimmune IgG-agarose column was analyzed by SDS-polacrylamide gel electrophoresis (PAGE). The material uniquely eluted from the immune agarose was represented by protein bands with apparent mol. wts. of 166,000, 130,000, and 76,000 and also retained some remnant antigen activity in antibody neutralization studies. Thrombin

activated factor VIII:C (40-fold) from von

willebrand factor-agarose chromatog. was also specifically bound only by the antifactor VIII:C-agarose. The inactive material which eluted from the antibody column contained polypeptides with apparent mol. wts. of 76,000, 67,000, and 50,000. Thus, the material purified by 2 different affinity reagents and visualized by the SDS-PAGE represents at least, in part, polypeptides derived from porcine

factor VIII:C.

L3 ANSWER 71 OF 72 MEDLINE on STN ACCESSION NUMBER: 82225663 MEDLINE DOCUMENT NUMBER: PubMed ID: 6806984

TITLE: Control of large-scale plasma thawing for recovery of

cryoprecipitate factor VIII.

AUTHOR: Foster P R; Dickson A J; McQuillan T A; Dickson I H; Keddie

S; Watt J G

SOURCE: Vox sanguinis, (1982) 42 (4) 180-9.

Journal code: 0413606. ISSN: 0042-9007.

PUB. COUNTRY: Switzerland

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 198208

ENTRY DATE: Entered STN: 19900317

Last Updated on STN: 19900317 Entered Medline: 19820826

AB Cryoprecipitation is commonly used as the primary step in the preparation of clinical factor VIII concentrates; yet recovery is usually very low. Much of this loss is due to poor temperature control and a process of continuous plasma thawing has been designed to overcome this. A substantial improvement has resulted, with an increase in both yield and purity of factor VIII:

C of over 50% in comparison to a conventional batch thaw process.

L3 ANSWER 72 OF 72 EMBASE COPYRIGHT 2005 ELSEVIER INC. ALL RIGHTS RESERVED.

on STN

ACCESSION NUMBER: 81209716 EMBASE

DOCUMENT NUMBER: 1981209716

TITLE: Kinetic analysis of bovine factor VIII in the hemophilic

dog.

AUTHOR: Gentry P.A.; Kirby E.P.; Gentry R.D.

CORPORATE SOURCE: Dept. Biomed. Scis Mathemat. Statist., Univ. Guelph,

Ontario, Canada

SOURCE: Thrombosis and Haemostasis, (1981) Vol. 46, No. 2, pp.

485-488.

CODEN: THHADQ

COUNTRY: Germany

DOCUMENT TYPE: Journal

FILE SEGMENT: 025 Hematology

LANGUAGE: English

ENTRY DATE: Entered STN: 911209

Last Updated on STN: 911209

AB Bovine factor VIII, which did not contain platelet aggregating factor activity, was infused into hemophilic dogs. Factor VIII procoagulant (VIII:C) levels in the dogs increased dramatically, then decreased in a biphasic manner. The half-life of the longest component was 3-7 hrs. The infusions were hemostatically effective and also caused a prolonged shortening of the activated partial thromboplastin time. These studies demonstrate that the platelet aggregating factor activity of bovine factor VIII is not essential for its maintenance in the circulation and that preparations lacking this activity may be clinically useful. When concentrates of partially purified factor VIII

:C (essentially free of both platelet aggregating factor and

:C (essentially free of both platelet aggregating factor and factor VIII-related antigen) were infused, marked increases in VIII:C levels were also observed, but the half-life was significantly shorter

(T1/2 of approximately 1 hr).

L6 ANSWER 1 OF 4 BIOTECHDS COPYRIGHT 2005 THE THOMSON CORP. on STN

ACCESSION NUMBER: 2004-14441 BIOTECHDS TITLE: Von Willebrand factor

von Willebrand lactor

concentrates containing Factor VIII:

c, having elevated high molecular multimer content

and low immunogenicity, useful for treating hemophilia A and

von Willebrand syndrome;

plasma or recombinant Factor-VIII:
c treatment for disease therapy

AUTHOR: KUMPE G; JURASCHEK M; MAYER N; SCHULTE S; WORMSBAECHER W

PATENT ASSIGNEE: AVENTIS BEHRING GMBH
PATENT INFO: EP 1405863 7 Apr 2004
APPLICATION INFO: EP 2003-20148 5 Sep 2003

PRIORITY INFO: DE 2002-1046125 1 Oct 2002; DE 2002-1046125 1 Oct 2002

DOCUMENT TYPE: Patent
LANGUAGE: German
AN 2004-14441 BIOTECHDS
AB DERWENT ABSTRACT:

NOVELTY - Von Willebrand factor (vWF)

concentrates (I) containing Factor VIII:C

(FVIII:C) are obtained by fractional precipitation

from a liquid containing FVIII:C and vWF; and have an elevated content of high molecular multimers of vWF and a ratio of vWF:RCoF (ristocetin cofactor) activity to vWF:Ag of more than 1, are new.

DETAILED DESCRIPTION - An INDEPENDENT CLAIM is included for the preparation of (I).

ACTIVITY - Hemostatic.

MECHANISM OF ACTION - Blood coagulation factor.

 \mbox{USE} - (I) are used in medicaments for treating hemophilia $\mbox{\sc A}$ and von Willebrand syndrome (claimed).

ADMINISTRATION - No details given in the source material.

ADVANTAGE - The high content of high molecular multimers of vWF reduces the immunogenicity of recombinant or plasma FVIII:C (claimed), and thus reduces side-effects. Optimal vWF/FVIII:C with enriched high molecular multimer content are obtainable in simple and targeted manner by selective precipitation. The coagulation-active FVIII:C is stabilized by the vWF, and the high content of high molecular multimers provides a more rapid hemostatic action.

EXAMPLE - A solution of 200 g cryoprecipitate in 800 ml 0.1 M sodium chloride-glycine solution was treated with 10 volume % of 1.5% aluminum hydroxide suspension, stirred for 15 minutes and centrifuged. The supernatant (820 ml) was stirred with sufficient glycine to precipitate the fibrinogen content and again centrifuged. The glycine-containing supernatant was treated under stirring with 15 % sodium chloride to precipitate the vWF/FVIII:C complex quantitatively. A solution of the precipitate in 64 ml sodium chloride-glycine buffer was stabilized with sucrose (1 g/ml) and glycine (150 g/l), pasteurized for 10 hours at 60 $\,$ degreesC, cooled and diluted with an equal volume of sodium chloride-glycine buffer. The diluted solution (containing 1.6 g/l sodium chloride and 124.4 g/l glycine) was treated under stirring with a precipitation medium such that the precipitation mixture contained 80 g/l glycine and 122 g/l sodium chloride. After stirring for 45 minutes, the fine precipitate was removed by centrifugation. The obtained fraction (dissolved in isotonic buffer) was enriched in high molecular multimers, and had ratios of FVIII:C to vWF:RCoF of 1:2.4, FVIII:C to vWF:Ag of 1:0.7 and vWF:Ag to vWF:RCoF of 1:3.6. For comparison, the starting complex had ratios of FVIII:C to vWF:RCoF of 1:3.1, FVIII:C to vWF:Ag of 1:2.5 and vWF:Ag to vWF:RCoF of 1:1.2; and cryo-solution after aluminum hydroxide adsorption had ratios of FVIII:C to vWF:RCoF of 1:1.6, FVIII:C to vWF:Ag of 1:2.6 and vWF:Ag to vWF:RCoF of 1:0.6.(14 pages)

L6 ANSWER 2 OF 4 IFIPAT COPYRIGHT 2005 IFI on STN
AN 10625429 IFIPAT; IFIUDB; IFICDB
TITLE: CONCENTRATE OF A FACTOR VIII:
C-CONTAINING VON WILLEBRAND

FACTOR AND THE PROCESS RELATING THERETO

Juraschek; Manfred, Weimar, DE

INVENTOR(S):

Kumpe; Gerhardt, Wetter, DE Mayer; Natascha, Marburg, DE Schulte; Stefan, Marburg, DE

Wormshabacher; Wilfried, Kirchhain, DE

PATENT ASSIGNEE(S):

Unassigned

AGENT:

Finnegan, Henderson, Farabow, ; Garrett & Dunner, L.L.P., 1300 I Street, N. W., Washington, DC,

20005-3315, US

NUMBER PK DATE ______ PATENT INFORMATION: US 2004132654 A1 20040708 APPLICATION INFORMATION: US 2003-670563 20030926

> NUMBER DATE _____ DE 2002-102461252 20021001 US 2004132654 20040708 -

PRIORITY APPLN. INFO.: FAMILY INFORMATION:

Utility

Patent Application - First Publication

FILE SEGMENT:

DOCUMENT TYPE:

CHEMICAL

APPLICATION

NUMBER OF CLAIMS:

18

The invention relates to a concentrate and a process for producing a

factor VIII:C-containing von Willebrand factor by fractional

precipitation from a liquid comprising factor

VIII: C and von Willebrand

factor, resulting in an increased content of high molecular

weight multimers of von Willebrand factor

and a ratio of the vWF:RCoF activity to vWF:Ag of greater than 1.

CLMN 18

TITLE:

ANSWER 3 OF 4 USPATFULL on STN 1.6

ACCESSION NUMBER:

\\\/2004:172490 USPATFULL

Concentrate of a factor VIII: C-containing von Willebrand

factor and the process relating thereto

INVENTOR(S):

Kumpe, Gerhardt, Wetter, GERMANY, FEDERAL REPUBLIC OF

Juraschek, Manfred, Weimar, GERMANY, FEDERAL REPUBLIC

Mayer, Natascha, Marburg, GERMANY, FEDERAL REPUBLIC OF Schulte, Stefan, Marburg, GERMANY, FEDERAL REPUBLIC OF Wormshabacher, Wilfried, Kirchhain, GERMANY, FEDERAL

REPUBLIC OF

NUMBER KIND DATE ______ US 2004132654 · A1 20040708 PATENT INFORMATION: US 2003-670563 APPLICATION INFO.: A1 20030926 (10)

NUMBER DATE _____ DE 2002-10246125 20021001

PRIORITY INFORMATION:

Utility

DOCUMENT TYPE: FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE:

Finnegan, Henderson, Farabow,, Garrett & Dunner,

L.L.P., 1300 I Street, N.W., Washington, DC, 20005-3315

NUMBER OF CLAIMS: 18

-1

EXEMPLARY CLAIM: LINE COUNT:

722

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The invention relates to a concentrate and a process for producing a

factor VIII: C-containing von Willebrand factor by fractional

precipitation from a liquid comprising factor

VIII: C and von Willebrand

factor, resulting in an increased content of high molecular
weight multimers of von Willebrand factor
and a ratio of the vWF:RCoF activity to vWF:Ag of greater than 1.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L6 ANSWER 4 OF 4 WPIDS COPYRIGHT 2005 THE THOMSON CORP on STN

ACCESSION NUMBER: 2004-297054 [28] WPIDS

DOC. NO. CPI: C2004-113576

TITLE: Von Willebrand factor

concentrates containing Factor VIII

c, having elevated high molecular multimer

content and low immunogenicity, useful for treating

hemophilia A and von Willebrand syndrome.

DERWENT CLASS: B04 D16

INVENTOR(S): JURASCHEK, M; KUMPE, G; MAYER, N; SCHULTE, S;

WORMSBAECHER, W; WORMSHABACHER, W

PATENT ASSIGNEE(S): (AVET) AVENTIS BEHRING GMBH; (CENT-N) CENTEON PHARMA

GMBH; (JURA-I) JURASCHEK M; (KUMP-I) KUMPE G; (MAYE-I) MAYER N; (SCHU-I) SCHULTE S; (WORM-I) WORMSHABACHER W

COUNTRY COUNT: 35

PATENT INFORMATION:

PATENT NO	KIND DATE	WEEK	LΑ	PG		
EP 1405863 R: AL AT BE		(200428) * (U IE IT	LI LT LU LV
MC MK NL	PT RO SE SI	SK TR				
CA 2443463	A1 20040401	. (200428) E	EN		•	
DE 10246125	A1 20040415	(200428)				
JP 2004123744	A 20040422	(200428)		16		
US 2004132654	A1 20040708	(200445)				
KR 2004030369	A 20040409	(200453)				

APPLICATION DETAILS:

PF	ATENT NO	KIND	APPLICATION	DATE
E	2 1405863	A1	EP 2003-20148	20030905
C.F	4 2443463	A1	CA 2003-2443463	20030929
DE	E 10246125	A1	DE 2002-10246125	20021001
JE	2004123744	A	JP 2003-339076	20030930
US	2004132654	A1	US 2003-670563	20030926
KF	R 2004030369	Α	KR 2003-68405	20031001

PRIORITY APPLN. INFO: DE 2002-10246125 20021001

AN 2004-297054 [28] WPIDS

AB EP 1405863 A UPAB: 20040429

NOVELTY - Von Willebrand factor (VWF)

concentrates (I) containing Factor VIII:C

(FVIII:C) are obtained by fractional precipitation .

from a liquid containing FVIII:C and vWF; and have an elevated content of high molecular multimers of vWF and a ratio of vWF:RCoF (ristocetin cofactor) activity to vWF:Ag of more than 1, are new.

DETAILED DESCRIPTION - An INDEPENDENT CLAIM is included for the preparation of (I).

ACTIVITY - Hemostatic.

MECHANISM OF ACTION - Blood coagulation factor.

 ${\tt USE-(I)}$ are used in medicaments for treating hemophilia A and von Willebrand syndrome (claimed).

ADVANTAGE - The high content of high molecular multimers of vWF reduces the immunogenicity of recombinant or plasma FVIII:C (claimed), and thus reduces side-effects. Optimal vWF/FVIII:C with enriched high molecular multimer content are obtainable in simple and targeted manner by selective precipitation. The coagulation-active FVIII:C is stabilized by the vWF, and the high content of high molecular multimers provides a more rapid hemostatic action.

```
=> d his
```

L3

L5

(FILE 'HOME' ENTERED AT 11:04:28 ON 02 AUG 2005)

FILE 'ADISCTI, ADISINSIGHT, ADISNEWS, AGRICOLA, ANABSTR, ANTE, AQUALINE, AQUASCI, BIOBUSINESS, BIOCOMMERCE, BIOENG, BIOSIS, BIOTECHDS, BIOTECHNO, CABA, CANCERLIT, CAPLUS, CEABA-VTB, CEN, CIN, CONFSCI, CROPB, CROPU, DDFB, DGENE, DISSABS, DRUGB, DRUGMONOG2, ...' ENTERED AT 11:05:21 ON 02 AUG 2005

L1 98 S FACTOR VIII:C AND WILLEBRAND FACTOR AND CONCENTRATE AND PREPA

L2 . 77 S L1 AND PY<2002

72 DUP REM L2 (5 DUPLICATES REMOVED)

L4 0 S L3 AND ALKALI METAL AND (FRACTIONAL PRECIPITATION)

0 S L3 AND FRACTIONAL PRECIPITATION

L6 4 S FRACTIONAL PRECIPITATION AND FACTOR VIII: C AND VON WILLEBRAND

=> s L6 and metal

40 FILES SEARCHED...

70 FILES SEARCHED...

L7 3 L6 AND METAL

=> d L7 1-3 ibib, abs

L7 ANSWER 1 OF 3 IFIPAT COPYRIGHT 2005 IFI on STN

AN 10625429 IFIPAT;IFIUDB;IFICDB

TITLE: C-CONTENTRATE OF A FACTOR VIII: C-CONTAINING VON WILLEBRAND

FACTOR AND THE PROCESS RELATING THERETO

INVENTOR(S): Juraschek; Manfred, Weimar, DE

Kumpe; Gerhardt, Wetter, DE Mayer; Natascha, Marburg, DE Schulte; Stefan, Marburg, DE

Wormshabacher; Wilfried, Kirchhain, DE

PATENT ASSIGNEE(S): Unassigned

AGENT: Finnegan, Henderson, Farabow,; Garrett & Dunner,

L.L.P., 1300 I Street, N. W., Washington, DC,

20005-3315, US

NUMBER DATE
----DE 2002-102461252 20021001
US 2004132654 20040708

FAMILY INFORMATION:
DOCUMENT TYPE:

PRIORITY APPLN. INFO.:

Utility

Patent Application - First Publication

FILE SEGMENT: CHEMICAL

APPLICATION

NUMBER OF CLAIMS: 18

AB The invention relates to a concentrate and a process for producing a

factor VIII:C-containing von
Willebrand factor by fractional

precipitation from a liquid comprising factor

VIII:C and von Willebrand

factor, resulting in an increased content of high molecular

weight multimers of von Willebrand factor

and a ratio of the vWF:RCoF activity to vWF:Ag of greater than 1.

CLMN 18

L7 ANSWER 2 OF 3 USPATFULL on STN

ACCESSION NUMBER: 2004:172490 USPATFULL

TITLE: Concentrate of a factor VIII:

C-containing von Willebrand

factor and the process relating thereto

INVENTOR(S): Kumpe, Gerhardt, Wetter, GERMANY, FEDERAL REPUBLIC OF

Juraschek, Manfred, Weimar, GERMANY, FEDERAL REPUBLIC

OF

Mayer, Natascha, Marburg, GERMANY, FEDERAL REPUBLIC OF Schulte, Stefan, Marburg, GERMANY, FEDERAL REPUBLIC OF Wormshabacher, Wilfried, Kirchhain, GERMANY, FEDERAL

REPUBLIC OF

NUMBER KIND DATE

PATENT INFORMATION: US 2004132654 A1 20040708

APPLICATION INFO.: US 2003-670563 A1 20030926 (10)

NUMBER DATE

PRIORITY INFORMATION: DE 2002-10246125 20021001

DOCUMENT TYPE: Utility FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: Finnegan, Henderson, Farabow,, Garrett & Dunner,

L.L.P., 1300 I Street, N.W., Washington, DC, 20005-3315

NUMBER OF CLAIMS: 18
EXEMPLARY CLAIM: 1
LINE COUNT: 722

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB The invention relates to a concentrate and a process for producing a

factor VIII:C-containing von
Willebrand factor by fractional

precipitation from a liquid comprising factor

VIII: C and von Willebrand

factor, resulting in an increased content of high molecular

weight multimers of von Willebrand factor

and a ratio of the vWF:RCoF activity to vWF:Ag of greater than 1.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L7 ANSWER 3 OF 3 WPIDS COPYRIGHT 2005 THE THOMSON CORP on STN

ACCESSION NUMBER: 2004-297054 [28] WPIDS

DOC. NO. CPI: C2004-113576

TITLE: Von Willebrand factor

concentrates containing Factor VIII

c, having elevated high molecular multimer

content and low immunogenicity, useful for treating

hemophilia A and von Willebrand syndrome.

DERWENT CLASS: B04 D16

INVENTOR(S): JURASCHEK, M; KUMPE, G; MAYER, N; SCHULTE, S;

WORMSBAECHER, W; WORMSHABACHER, W

PATENT ASSIGNEE(S): (AVET) AVENTIS BEHRING GMBH; (CENT-N) CENTEON PHARMA

GMBH; (JURA-I) JURASCHEK M; (KUMP-I) KUMPE G; (MAYE-I)

MAYER N; (SCHU-I) SCHULTE S; (WORM-I) WORMSHABACHER W .

COUNTRY COUNT: 35

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA PG

EP 1405863 A1 20040407 (200428)* GE 14

R: AL AT BE BG CH CY CZ DE DK EE ES FI FR GB GR HU IE IT LI LT LU LV

MC MK NL PT RO SE SI SK TR

CA 2443463 A1 20040401 (200428) EN

DE 10246125 A1 20040415 (200428)

JP 2004123744 A 20040422 (200428) 16

US 2004132654 A1 20040708 (200445)

KR 2004030369 A 20040409 (200453)

APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
EP 1405863	Al	EP 2003-20148	20030905
CA 2443463	A1	CA 2003-2443463	20030929
DE 10246125	A1	DE 2002-10246125	20021001
JP 2004123744	A	JP 2003-339076	20030930
US 2004132654	A1	US 2003-670563	20030926
KR 2004030369	A	KR 2003-68405	20031001

PRIORITY APPLN. INFO: DE 2002-10246125 20021001

AN 2004-297054 [28] WPIDS

EP 1405863 A UPAB: 20040429

NOVELTY - Von Willebrand factor (vWF)

concentrates (I) containing Factor VIII:C

(FVIII:C) are obtained by fractional precipitation

from a liquid containing FVIII:C and vWF; and have an elevated content of high molecular multimers of vWF and a ratio of vWF:RCoF (ristocetin cofactor) activity to vWF:Ag of more than 1, are new.

DETAILED DESCRIPTION - An INDEPENDENT CLAIM is included for the preparation of (I).

ACTIVITY - Hemostatic.

MECHANISM OF ACTION - Blood coagulation factor.

 \mbox{USE} - (I) are used in medicaments for treating hemophilia A and von Willebrand syndrome (claimed).

ADVANTAGE - The high content of high molecular multimers of vWF reduces the immunogenicity of recombinant or plasma FVIII:C (claimed), and thus reduces side-effects. Optimal vWF/FVIII:C with enriched high molecular multimer content are obtainable in simple and targeted manner by selective precipitation. The coagulation-active FVIII:C is stabilized by the vWF, and the high content of high molecular multimers provides a more rapid hemostatic action. Dwg.0/0

=> d his

AB

(FILE 'HOME' ENTERED AT 11:04:28 ON 02 AUG 2005)

FILE 'ADISCTI, ADISINSIGHT, ADISNEWS, AGRICOLA, ANABSTR, ANTE, AQUALINE, AQUASCI, BIOBUSINESS, BIOCOMMERCE, BIOENG, BIOSIS, BIOTECHDS, BIOTECHNO, CABA, CANCERLIT, CAPLUS, CEABA-VTB, CEN, CIN, CONFSCI, CROPB, CROPU, DDFB, DGENE, DISSABS, DRUGB, DRUGMONOG2, ...' ENTERED AT 11:05:21 ON 02 AUG 2005

- 98 S FACTOR VIII: C AND WILLEBRAND FACTOR AND CONCENTRATE AND PREPA
- 77 S L1 AND PY<2002
 - 72 DUP REM L2 (5 DUPLICATES REMOVED)
 - O S L3 AND ALKALI METAL AND (FRACTIONAL PRECIPITATION)
 - 0 S L3 AND FRACTIONAL PRECIPITATION
 - 4 S FRACTIONAL PRECIPITATION AND FACTOR VIII:C AND VON WILLEBRAND
- 3 S L6 AND METAL

L1

L2

L3

L4 L5

L6

L7